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(54) Title: PLANT GENES FOR SENSITIVITY TO ETHYLENE AND PATHOGENS (57) Abstract <p>The present invention is directed to nucleic acid sequences for ethylene insensitive, EIN loci and corresponding amino acid sequences. The present invention is also directed to nucleic acid sequences for hookless 1, HLS1, alleles and amino acid sequences.</p>			

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PLANT GENES FOR SENSITIVITY TO ETHYLENE AND PATHOGENS

REFERENCE TO RELATED APPLICATIONS

This application is a continuation-in-part of U.S. application Serial No. 08/003,311, filed January 12, 1993, a continuation-in-part of U.S. application Serial No. 928,464, filed August 10, 1992; this application is also a continuation-in-part of U.S. application Serial No. 08/171,207, filed December 21, 1993, which is a continuation of U.S. application Serial No. 899,262, filed June 16, 1992, now abandoned; the disclosures of which are hereby incorporated in their entirety.

REFERENCE TO GOVERNMENT GRANTS

This work was supported in part by research grants from the National Institutes of Health GM-26379 and National Science Foundation grant IBN-92-05342. The United States Government may have certain rights in this invention.

BACKGROUND OF THE INVENTION

Ethylene, a gaseous plant hormone, is involved in the regulation of a number of plant processes ranging from growth and development to fruit ripening. As in animal systems, response of plants to disease not only involves static processes, but also involves inducible defense mechanisms. One of the earliest detectable event to occur during plant-pathogen interaction is a rapid increase in ethylene biosynthesis. Ethylene biosynthesis, in response to pathogen invasion, correlates with increased defense

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mechanisms, chlorosis, senescence and abscission. The molecular mechanisms underlying operation of ethylene action, however, are unknown. Nonetheless, ethylene produced in response to biological stress is known to regulate the rate of transcription of specific plant genes. A variety of biological stresses can induce ethylene production in plants including wounding, bacterial, viral or fungal infection as can treatment with elicitors, such as glycopeptide elicitor preparations (prepared by chemical extraction from fungal pathogen cells). Researchers have found, for example, that treatment of plants with ethylene generally increases the level of many pathogen-inducible "defense proteins", including β -1,3-glucanase, chitinase, L-phenylalanine ammonia lyase, and hydroxyproline-rich glycoproteins. The genes for these proteins can be transcriptionally activated by ethylene and their expression can be blocked by inhibitors of ethylene biosynthesis. Researchers have also characterized a normal plant response to the production or administration of ethylene, as a so-called "triple response". The triple response involves inhibition of root and stem elongation, radial swelling of the stem and absence of normal geotropic response (diageotropism).

Ethylene is one of five well-established plant hormones. It mediates a diverse array of plant responses including fruit ripening, leaf abscission and flower senescence.

The pathway for ethylene biosynthesis has been established (Figure 6). Methionine is converted to ethylene with S-adenylmethionine (SAM) and 1-aminocyclopropane-1-carboxylic acid (ACC) as intermediates. The production of ACC from SAM is catalyzed by the enzyme ACC synthase. Physiological analysis has suggested that this is the key regulatory step in the pathway, see Kende, *Plant Physiol.* 1989, 91, 1-4. This enzyme has been cloned from several sources, see Sato et al., *PNAS, (USA)* 1989, 86, 6621; Van Der Straeten et al.,

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PNAS, (USA) 1990, 87, 4859-4863; Nakajima et al., *Plant Cell Physiol.* 1990, 29, 989. The conversion of ACC to ethylene is catalyzed by ethylene forming enzyme (EFE), which has been recently cloned (Spanu et al., *EMBO J* 1991, 10, 2007. Aminoethoxy-vinylglycine (AVG) and α -aminoisobutyric acid (AIB) have been shown to inhibit ACC synthase and EFE respectively. Ethylene binding is inhibited non-competitively by silver, and competitively by several compounds, the most effective of which is trans-cyclooctane. ACC synthase is encoded by a highly divergent gene family in tomato and *Arabidopsis* (Theologis, A., *Cell* 70:181 (1992)). ACC oxidase, which converts ACC to ethylene, is expressed constitutively in most tissues (Yang et al., *Ann. Rev. Plant Physiol.* 1984, 35, 155), but is induced during fruit ripening (Gray et al. *Cell* 1993 72, 427). It has been shown to be a dioxygenase belonging to the Fe²⁺/ascorbate oxidase superfamily (McGarvey et al., *Plant Physiol.* 1992, 98, 554).

Etiolated dicotyledonous seedlings are normally highly elongated and display an apical arch-shaped structure at the terminal part of the shoot axis; the apical hook. The effect of ethylene on dark grown seedlings, the triple response, was first described in peas by Neljubow in 1901, Neljubow, D., *Pflanzen Beih. Bot. Zentralb.*, 1901, 10, 128. In *Arabidopsis*, a typical triple response consists of a shortening and radial swelling of the hypocotyl, an inhibition of root elongation and an exaggeration of the curvature of the apical hook (Figures 7 and 16). Etiolated morphology is dramatically altered by stress conditions which induce ethylene production the ethylene-induced "triple response" may provide the seedling with additional strength required for penetration of compact soils, see Harpham et al., *Annals of Bot.*, 1991, 68, 55. Ethylene may also be important for other stress responses. ACC synthase gene expression and ethylene production is induced by many types of biological and physical stress, such as wounding and pathogen infection,

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see Boller, T., in *The Plant Hormone Ethylene*, A.K. Mattoo and J.C. Suttle eds., 293-314, 1991, CRC Press, Inc. Boca Raton and Yu, Y. et al., *Plant Phys.*, 1979, 63, 589, Abeles et al. 1992 Second Edition San Diego, CA Academic Press;
5 and Gray et al. *Plant Mol Biol.* 1992 19, 69.

A number of researchers have identified the interaction between *Arabidopsis thaliana* and *Pseudomonas syringae* bacteria; Whalen et al., "Identification of *Pseudomonas syringae* Pathogens of *Arabidopsis* and a
10 Bacterial Locus Determining Avirulence on Both *Arabidopsis* and Soybean", *The Plant Cell* 1991, 3, 49, Dong et al., "Induction of *Arabidopsis* Defense Genes by Virulent and Avirulent *Pseudomonas syringae* Strains and by a Cloned Avirulence Gene", *The Plant Cell* 1991, 3, 61, and Debener
15 et al., "Identification and Molecular Mapping of a Single *Arabidopsis thaliana* Locus Determining Resistance to a Phytopathogenic *Pseudomonas syringae* Isolate", *The Plant Journal* 1991, 1, 289. *P. syringae* pv. tomato (Pst) strains are pathogenic on *Arabidopsis*. A single bacterial gene,
20 avrRpt2, was isolated that controls pathogen avirulence on specific *Arabidopsis* host genotype Col-0.

Bent, A.F., et al., "Disease Development in Ethylene-Insensitive *Arabidopsis thaliana* Infected with Virulent and Avirulent *Pseudomonas* and *Xanthomonas*
25 Pathogens", *Molecular Plant-Microbe Interactions* 1992, 5, 372; Agrios, G.N., *Plant Pathology* 1988, 126, Academic Press, San Diego; and Mussel, H., "Tolerance to Disease", page 40, in *Plant Disease: An Advanced Treatise, Volume 5*, Horsfall, J.G. and Cowling, E.B., eds., 1980, Academic
30 Press, New York, establish the art recognized definitions of tolerance, susceptibility, and resistance. Tolerance is defined for purposes of the present invention as growth of a pathogen in a plant where the plant does not sustain damage. Resistance is defined as the inability of a
35 pathogen to grow in a plant and no damage to the plant results. Susceptibility is indicated by pathogen growth with plant damage.

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Regardless of the molecular mechanisms involved, the normal ethylene response of a plant to pathogen invasion has been thought to have a cause and effect relationship in the ability of a plant to fight off plant pathogens. Plants insensitive in any fashion to ethylene were believed to be incapable of eliciting a proper defense response to pathogen invasion, and thus unable to initiate proper defense mechanisms. As such, ethylene insensitive plants were thought to be less disease tolerant.

10 The induction of disease responses in plants requires recognition of pathogens or pathogen-induced symptoms. In a large number of plant-pathogen interactions, successful resistance is observed when the plant has a resistance gene with functional specificity for pathogens that carry a particular avirulence gene. If the plant and pathogen carry resistance and avirulence genes with matched specificity, disease spread is curtailed and a hypersensitive response involving localized cell death and physical isolation of the pathogen typically occurs. In 15 the absence of matched resistance and avirulence genes, colonization and tissue damage proceed past the site of initial infection and disease is observed.

A better understanding of plant pathogen tolerance is needed. Also needed is the development of methods for improving the tolerance of plants to pathogens, as well as the development of easy and efficient methods for identifying pathogen tolerant plants.

Genetic and molecular characterization of several gene loci and protein products is set forth in the present invention. The results will reveal interactions among modulatory components of the ethylene action pathway and provide insight into how plant hormones function. Thus, the quantity, quality and longevity of food, such as fruits and vegetables, and other plant products such as flowers, 30 will be improved thereby providing more products for market in both developed and underdeveloped countries.

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SUMMARY OF THE INVENTION

The present invention is directed to nucleic acid sequences for ethylene insensitive, EIN loci and corresponding amino acid sequences. Several ein wild type sequences, mutations, amino acid sequences, and protein products are included within the scope of the present invention. The nucleic acid sequences set forth in SEQUENCE ID NUMBERS 1 and 2 for ein2; 4, 5, 7, 9, and 11 for ein3 and eil1, eil2, eil3; as well as amino acid sequences set forth in SEQUENCE ID NUMBERS 3 for ein2; 6, 8, 10, 12, and 13 for ein3 and eil1, eil2, eil3; are particular embodiments of the present invention.

The present invention is also directed to nucleic acid sequences for hookless1, HLS1, alleles and amino acid sequences. Wild type and mutated nucleic acid sequences, amino acid sequences and proteins are included within the scope of the present invention. The nucleic acid sequences of hls1 are set forth in SEQUENCE ID NUMBERS: 14 and 15; the amino acid sequences are set forth in SEQUENCE ID NUMBER: 16.

These and other aspects of the invention will become more apparent from the following detailed description when taken in conjunction with the following figures.

BRIEF DESCRIPTION OF THE FIGURES

Figure 1 displays the EIN2 region on chromosome 5 of *Arabidopsis thaliana*. ○ represents the left end probe, □ represents the right end probe, a length of 100 kb is represented in the legend.

Figure 2 is a genomic Southern blot. A polymorphism was detected in ein2-12 by hybridization with g3715. The g3715 cosmid was hybridized to a genomic Southern blot containing several alleles of ein2. In ein2-12 EcoR I digested genomic DNA, two bands were missing, 1.2 kb and 4.3 kb; and a new 5.5 kb fragment was detected. The DNA from the ein2 alleles was purified according to Chang et al. Proc. Natl. Acad. Sci USA 1988 85, 6857. 5 µg of

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EcoR I digested DNA was separated on a 0.8% agarose gel and blotted to hybrid N⁺ (Sambrook et al., *Molecular Cloning: A Laboratory Manual*, 2nd ed., 1989, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, Amersham, Arlington Heights, IL). All hybridizations were done using random hexamer labeled DNAs (Feinberg and Volgelstein, *Anal. Biochem* 1984 137, 266). Filters were prehybridized for at least 2 hours in 0.5 M sodium phosphate pH 7.2, 7% sodium dodecyl sulfate, and 1% BSA at 60° C. Hybridization of a minimum of 15 hours was in a solution of 0.5 M sodium phosphate pH 7.2, 7% sodium dodecyl sulfate, and 1% BSA at 60° C. Hybridization filters were washed and autoradiographed (Sambrook et al. 1989).

Figure 3 is a diagram of the polymorphism in *ein2-12* due to the loss of an *EcoR I* site. The pgEE1.2 subclone from g3715 is shown.

Figure 4 is a description of the *EIN2* locus, the cDNA (bottom) is shown relative to the genomic map (top). A putative TATA sequence is shown approximately 60 base pairs 5' to the start of the cDNA. The position of the translation start and stop sites are also shown.

Figure 5 exhibits the sequence of the *EIN2* locus. Genomic DNA sequence (SEQUENCE ID NO: 1) is shown in lower case letters, cDNA sequence (SEQUENCE ID NO: 2) is shown in capital letters. The predicted peptide sequence (SEQUENCE ID NO: 3) is displayed under the corresponding nucleic acid codons.

Figure 6 is a schematic illustration of the ethylene biosynthesis pathway.

Figure 7 depicts a seedling body and developing plant. Specifically, Figure 7A is a cross section of the seedling body of a seed plant. Figure 7B is a perspective view of a developing seed plant.

Figure 8 identifies the protein sequences of *eil1*, *ein3*, *eil2*, *eil3*, and a common consensus protein sequence representing all four of the individual protein sequences.

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Figure 9 displays the *EIN3* gene structure and mutants. Also set forth in Figure 9 is the predicted polypeptide acidity and basicity, as well as Asn repeats.

Figure 10 exhibits a map of chromosome 3 and the position of *EIN3* relative to other gene loci.

Figure 11 sets forth a map of chromosome 2 and the position of *EIL1* relative to other gene loci.

Figure 12 displays a map of chromosome 5 and the position of *EIL2* relative to other gene loci.

Figure 13 exhibits a map of chromosome 4 and the position of *HLS1* relative to other gene loci.

Figure 14 is a representation of the arrangement of *hls* mutants on chromosome 4.

Figure 15 identifies the protein sequences of *Arabidopsis HLS1* and acetyl transferases in *E. coli*, *Pseudomonas*, *Streptomyces*, Mouse, Human, *Azospirillum*, Yeast, and *Citrobacter*. A consensus sequence representing common amino acids of the sequences is also provided.

Figure 16 displays ethylene responses in wild type and mutant: *ctrl1*, *etol1*, *hls1*, *etr1*, *ein2*, *ein3*, *Arabidopsis* seedlings. Seeds of the indicated genotype were germinated and grown for three days in the dark in either air or air containing 10 ppm ethylene.

Figure 17 is a genetic model of interactions among components of the ethylene signal transduction pathway. This model shows the predicted order in which the various gene products act which is based on the epistatic relationships among the mutants. The seedling ethylene responses are indicated on the right.

Figure 18 is a representation of pNLEIN3Bg12 indicating the relationship between the promoter, GUS, and *EIN3* sequences.

Figure 19 displays *EIN3* sequences. Figure 19A sets forth *EIN3* cDNA (SEQUENCE ID NO: 4), Figure 19B sets forth *EIN3* genomic DNA (SEQUENCE ID NO: 5), and Figure 19C sets forth *EIN3* protein sequence (SEQUENCE ID NO: 6).

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Figure 20 displays EIL1 sequences. Figure 20A sets forth EIL1 cDNA (SEQUENCE ID NO: 7), Figure 20B sets forth EIL1 peptide sequence (SEQUENCE ID NO: 8).

Figure 21 displays EIL2 sequences. Figure 21A sets forth EIL2 cDNA (SEQUENCE ID NO: 9), Figure 21B sets forth EIL2 peptide sequence (SEQUENCE ID NO: 10).

Figure 22 displays EIL3 sequences. Figure 22A sets forth EIL3 cDNA (SEQUENCE ID NO: 11). EIL3 peptide sequence is set forth in SEQUENCE ID NO: 12.

Figure 23 displays HLS1 sequences. Figure 23A sets forth HLS1 cDNA (SEQUENCE ID NO: 14), Figure 23B sets forth HLS1 genomic DNA sequence (SEQUENCE ID NO: 15), and Figure 23C sets forth HLS1 peptide sequence.

DETAILED DESCRIPTION OF THE INVENTION

The present invention is directed to nucleic acid and amino acid sequences which lend valuable characteristics to plants.

The present invention is directed to nucleic acid sequences of the *EIN2* locus. Wild type and mutant sequences of *EIN2* are within the scope of the present invention. Amino acid and protein sequences corresponding to the nucleic acid sequences are included in the present invention. *EIN2* mutations provide for ethylene insensitivity and pathogen tolerance in plants.

SEQUENCE ID NO: 2, the isolated cDNA representing the nucleic acid sequence coding for *EIN2* and the isolated genomic *EIN2* sequence of SEQUENCE ID NO: 1 are embodiments of the present invention. The purified amino acid sequence of SEQUENCE ID NO: 3 represents the *EIN2* protein product encoded by the cDNA identified above. The *EIN2* mutations identified herein by nucleotide position are measured in accordance with the beginning of the cDNA.

An *ein2-3* mutation was created by X-ray mutagenesis which resulted in a thymidine insertion at nucleotide position 3642 of the cDNA sequence in SEQUENCE

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ID NO: 2. A frameshift results in the corresponding amino acid sequence.

An *ein2-4* mutation was also generated by X-ray mutagenesis. The *ein2-4* mutation has an "AG" to "TTT" mutation at position 2103 of the *EIN2* cDNA sequence resulting in a frameshift in the corresponding amino acid sequence.

An *ein2-5* mutation was generated by X-ray mutagenesis, such that a deletion beginning at nucleic acid position 1570 of the cDNA occurred. Nucleic acids CATGACT were deleted. A frameshift results in the corresponding protein product.

An *ein2-6* mutation has a deletion of nucleic acids GAGTTGCGCATG, SEQ ID NO: 17, beginning at nucleic acid position 965 of the cDNA sequence. The *ein2-6* mutation was generated by *Agrobacterium* mutagenesis. This mutation results in a deletion at the amino acid level of Gly-Val-Ala-His, SEQ ID NO: 18, formerly beginning at amino acid position 115.

Another mutation, *ein2-9* was generated by DEB mutagenesis and has an "A" to "C" transition at position 4048 that results in a "His" to "Pro" change at amino acid position 1143 in the corresponding protein.

ein2-11 was generated by DEB mutagenesis and has a "TG" to "AT" transition at nucleic acid position 3492. This results in an Ochre stop signal at amino acid position 957 in the protein.

An *ein2-12* mutation was obtained by X-ray mutagenesis resulting in a deletion at nucleic acid position 1611 of nucleic acids TGCTACAATCAGAATTCTTGCACT, SEQ ID NO: 19. The corresponding amino acid sequence reveals a deletion of amino acids Ala-Thr-Ile-Arg-Ile-Leu-Ala-Val, SEQ ID NO: 20, beginning at amino acid position 331.

An *ein2-16* mutation results in an "AGT" to "G" transition at nucleic acid position 2851 as a result of X-

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ray mutagenesis. A frameshift results in the corresponding protein.

Table 4 sets forth the *EIN2* alleles and the results of the mutagenesis.

5 *Ein3* sequences for genes and proteins are the subject of the present invention. The present invention is directed to wild type nucleic acid and amino acid sequences as well as mutations of these sequences. *EIN3* mutations result in ethylene insensitive plants. *Ein*-like genes and
10 protein sequences, including *eil1*, *eil2*, and *eil3* sequences, are similar to *ein3* sequences, and are also disclosed in the present invention. The *EIN3* mutations are identified below by nucleotide position number in accordance with the beginning of the genomic DNA sequence.

15 The DNA sequences coding for *ein3* are set forth in SEQ ID NOS: 5 (genomic) and 4 (cDNA). The amino acid sequence may be found in SEQ ID NO: 6.

 In *ein3-1*, a "G" to "A" conversion in the genomic DNA at nucleotide 1598 occurs as a result of EMS
20 mutagenesis. In the corresponding protein, "W" is changed to a stop codon at amino acid position 215. The *ein3-2* mutation was generated by T-DNA insertion mutagenesis. The T-DNA inserted after nucleotide 2001 of the genomic, interrupting the protein after amino acid 349. The *ein3-3*
25 mutation results in a "G" to "T" switch at nucleotide position 1688 of genomic DNA as a result of DEB mutagenesis. The amino acid sequence results in a conversion of "K" to "N" at amino acid position 245.

 The cDNAs of *eil1*, *eil2*, and *eil3*, are set forth
30 in SEQ ID NOS: 7, 9, and 11, respectively. The corresponding amino acid sequences for the *ein*-like genes are set forth in SEQ ID NOS: 8, 10, and 12, (*eil1*, *eil2*, and *eil3*, respectively). A consensus sequence representing the common codons of the three *ein*-like genes is SEQ ID NO:
35 13.

Table 6 sets forth the *EIN3* alleles and the results of the mutagenesis. The translation start site of

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EIN3 is at nucleotide position 954 of the genomic sequence. the translation start sites for EIL1, EIL2, and EIL3 are at nucleotide positions 251, 8, and 102 of the respective cDNA sequences.

5 The present invention is directed to wild type and mutant sequences for the *Hls1* locus. The *hls* gene is regulated by ethylene directly. Amino acid and protein sequences corresponding to the wild type and mutant gene for *Hls1* are within the scope of the present invention.

10 The present invention is directed to nucleic acid sequences of the *HLS1* locus. Wild type and mutant sequences of *HLS1* are within the scope of the present invention. Amino acid and protein sequences corresponding to the nucleic acid sequences are included in the present
15 invention. The *HLS1* mutations are identified below by nucleotide position number in accordance with the beginning of the genomic DNA sequence.

SEQUENCE ID NO: 14, the isolated cDNA representing the nucleic acid sequence coding for *HLS1*, and
20 the isolated genomic *HLS1* sequence of SEQUENCE ID NO: 15 are embodiments of the present invention. The purified amino acid sequence of SEQUENCE ID NO: 16 represents the *HLS1* protein product encoded by the cDNA identified above.

An *hls1-1* mutation was created by EMS mutagenesis
25 which resulted in a "G" to "A" transition at nucleotide position 3487 of the genomic DNA sequence. This frameshift results in the corresponding amino acid sequence having a "Glu" to "Lys" substitution at amino acid position 345.

An *hls1-5* mutation of was generated by DEB
30 mutagenesis. The *hls1-5* mutation has an "T" to "A" mutation at position 2194 of the *HLS1* genomic DNA sequence, resulting in a mutation in the splice donor site. An *hls1-7* mutation was also created by DEB and resulted in a "T" to "A" transition at nucleic acid position 2194. The result
35 in the amino acid sequence is also a mutation in the splice donor site. Mutations at splice donor sites often result in aberrant splicing causing a frameshift or insertion to

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occur. The exact nature of the change in *hls1-5* and *hls1-7* may be determined by analyzing the protein from those mutants using an antibody.

hls1-6 is a mutation created by EMS resulting in
5 a "T" to "G" transition at nucleic acid position 3431. The corresponding amino acid sequence has a "Lys" to "Trp" substitution at amino acid position 326.

The mutation *hls1-4* was created by DEB
mutagenesis resulting in a "G" to "A" transition at nucleic
10 acid position 3487. The corresponding amino acid sequence has a "Glu" to "Lys" change at amino acid position 345.

hls1-9 is created by EMS mutagenesis. The
sequence results in "C" to "T" at nucleic acid position
2060, which corresponds to an "Arg" to "TGA" creating a
15 "stop signal" at amino acid position 11.

hls1-8 is a mutation resulting from EMS
mutagenesis. The nucleic acid sequence has a "C" to "T"
change at position 2992. The mutation results in an amino
acid sequence having an "Arg" to "Stop" transition at amino
20 acid position 180.

An EMS mutation resulting in a "G" to "A" change
at nucleic acid position 2033 is represented by *hls1-10*.
The amino acid sequence corresponding to the mutation
reveals a "Met" (Start signal) to "Ile" transition at amino
25 acid position 1.

Table 7 sets forth the *HLS1* alleles and the
results of the mutagenesis.

In accordance with the present invention, nucleic
acid sequences include and are not limited to DNA,
30 including and not limited to cDNA and genomic DNA; RNA,
including and not limited to mRNA and tRNA; and suitable
nucleic acid sequences such as those set forth in SEQUENCE
ID NUMBERS set forth herein, and alterations in the nucleic
acid sequences including alterations, deletions, mutations
35 and homologs. In addition, mismatches within the sequences
identified above, which achieve the methods of the
invention, are also considered within the scope of the

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disclosure. The sequences may also be unmodified or modified.

Also amino acid, peptide and protein sequences within the scope of the present invention include, and are
5 not limited to, the sequences set forth herein and alterations in the amino acid sequences including alterations, deletions, mutations and homologs.

In accordance with the invention, the nucleic acid sequences employed in the invention may be
10 exogenous/heterologous sequences. Exogenous and heterologous, as used herein, denotes a nucleic acid sequence which is not obtained from and would not normally form a part of the genetic make-up of the plant or the cell to be transformed, in its untransformed state. Plants
15 comprising exogenous nucleic acid sequences of *ein2*, *ein3*, *eil1*, *eil2*, *eil3*, or *hls1* mutations, such as and not limited to the nucleic acid sequences of SEQUENCE ID NUMBERS set forth herein are within the scope of the invention.

20 Transfected and/or transformed plant cells comprising nucleic acid sequences of *ein2*, *ein3*, *eil1*, *eil2*, *eil3*, or *hls1* mutations, such as and not limited to the nucleic acid sequences of SEQUENCE ID NUMBERS set forth herein, are within the scope of the invention. Transfected
25 cells of the invention may be prepared by employing standard transfection techniques and procedures as set forth in Sambrook et al., *Molecular Cloning: A Laboratory Manual*, 2nd ed., 1989, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, hereby incorporated by reference in
30 its entirety.

In accordance with the present invention, mutant plants which may be created with the sequences of the claimed invention include higher and lower plants in the Plant Kingdom. Mature plants and seedlings are included in
35 the scope of the invention. A mature plant includes a plant at any stage in development beyond the seedling. A

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seedling is a very young, immature plant in the early stages of development.

Particularly preferred plants are those from:
the Family Umbelliferae, particularly of the genera *Daucus*
5 (particularly the species *carota*, carrot) and *Apium*
(particularly the species *graveolens dulce*, celery) and the
like; the Family Solanaceae, particularly of the genus
Lycopersicon, particularly the species *esculentum* (tomato)
and the genus *Solanum*, particularly the species *tuberosum*
10 (potato) and *melongena* (eggplant), and the like, and the
genus *Capsicum*, particularly the species *annuum* (pepper) and
the like; and the Family Leguminosae, particularly the
genus *Glycine*, particularly the species *max* (soybean) and
the like; and the Family Cruciferae, particularly of the
15 genus *Brassica*, particularly the species *campestris*
(turnip), *oleracea* cv *Tastie* (cabbage), *oleracea* cv
Snowball Y (cauliflower) and *oleracea* cv *Emperor* (broccoli)
and the like; the Family Compositae, particularly the genus
Lactuca, and the species *sativa* (lettuce), and the genus
20 *Arabidopsis*, particularly the species *thaliana* (Thale
cress) and the like. Of these Families, the most preferred
are the leafy vegetables, for example, the Family
Cruciferae, especially the genus *Arabidopsis*, most
especially the species *thaliana*.

25 *Ein2* mutant sequences render plants disease and
pathogen tolerant, and ethylene insensitive. For purposes
of the current invention, disease tolerance is the ability
of a plant to survive infection with minimal injury or
reduction in the harvested yield of saleable material.
30 Plants with disease tolerance may have extensive levels of
infection but have little necrosis and few to no lesions.
These plants may also have reduced necrotic and water
soaking responses and chlorophyll loss may be virtually
absent. In contrast, resistant plants generally limit the
35 growth of pathogens and contain the infection to a
localized area with multiple apparent injurious lesions.

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The current invention is directed to, for example, identifying plant tolerance to bacterial infections including, but not limited to *Clavibacter michiganense* (formerly *Corynebacterium michiganense*),
5 *Pseudomonas solanacearum* and *Erwinia stewartii*, and more particularly, *Xanthomonas campestris* (specifically pathovars *campestris* and *vesicatoria*), *Pseudomonas syringae* (specifically pathovars *tomato*, *maculicola*).

In addition to bacterial infections, disease
10 tolerance to infection by other plant pathogens is within the scope of the invention. Examples of viral and fungal pathogens include, but are not limited to tobacco mosaic virus, cauliflower mosaic virus, turnip crinkle virus, turnip yellow mosaic virus; fungi including *Phytophthora*
15 *infestans*, *Peronospora parasitica*, *Rhizoctonia solani*, *Botrytis cinerea*, *Phoma lingam* (*Leptosphaeria maculans*), and *Albugo candida*.

Like *ein2*, *ein3* mutants also exhibit ethylene insensitivity. However, *ein3* mutants do not exhibit
20 disease or pathogen tolerance. Ethylene, $\text{CH}_2=\text{CH}_2$, is a naturally occurring plant hormone. The ethylene regulatory pathway includes the ethylene biosynthesis pathway and the ethylene autoregulatory or feedback pathway, see Figure 6. In the ethylene biosynthesis pathway, methionine is
25 converted to ethylene with S-adenosylmethionine (SAM) and 1-aminocyclopropane-1-carboxylic acid (ACC) as intermediates. These two reactions are catalyzed by ACC synthase and ethylene-forming enzyme (EFE), respectively. Little is known about the enzymes catalyzing these
30 reactions and their regulation at the molecular level.

The receptor and receptor complex of Figure 6 are believed to function with the autoregulatory pathway in the control of ethylene production. Ethylene regulatory
pathway inhibitors are positioned along the left side of
35 Figure 6. The inhibitors include AVG (aminoethoxyvinylglycine) and AIB (α -aminoisobutyric acid). The steps at which the mutants, ethylene overproducer (*etol*), ethylene

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insensitive (ein1, ein2) and hookless (hls1), are defective appear on the right of Figure 6.

In accordance with the claimed invention, ethylene insensitive plants are those which are unable to display a typical ethylene response when treated with high concentrations of ethylene. For purposes of the present invention, ethylene insensitivity includes total or partial inability to display a typical ethylene response. A typical ethylene response in wild type plants includes, for example, the so-called "triple response" which involves inhibition of root and stem elongation, radial swelling of the stem, and absence of normal geotropic response (diageotropism). Thus, for example, ethylene insensitive plants may be created in accordance with the present invention by the presence of an altered "triple response" wherein the root and stem are elongated despite the presence of high concentrations of ethylene. Further, a typical ethylene response also includes a shut down or diminution of endogenous ethylene production, upon application of high concentrations of ethylene. Ethylene insensitive plants may thus also be screened for, in accordance with the present invention, by the ability to continue production of ethylene, despite administration of high concentrations of ethylene. Such ethylene insensitive plants are believed to have impaired receptor function such that ethylene is constitutively produced despite the presence of an abundance of exogenous ethylene.

Screening includes screening for root or stem elongation and screening for increased ethylene production. Ethylene sensitive wild type plants experience an inhibition of root and stem elongation when an inhibitory amount of ethylene is administered. By inhibition of root and stem elongation, it is meant that the roots and stems grow less than the normal state (that is, growth without application of an inhibitory amount of ethylene). Typically, normal *Arabidopsis* (Col) grown without ethylene or ethylene precursor aminocyclopropane, ACC, root

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elongation is about 6.5 ± 0.2 mm/3 days; normal stem elongation is 8.7 ± 0.3 mm/3 days. Ein 2-1 plants grown without ethylene or ACC have root elongation of about 7.5 ± 0.2 mm/3 days and stem elongation of 11.35 ± 0.3 mm/3 days.

5 In the presence of $100 \mu\text{m}$ ACC, Col root growth is 1.5 ± 0.04 mm/3 days; ein 2-1 is 4.11 ± 0.1 mm/3 days and stem growth of 3.2 ± 0.1 mm/3 days for Col and 8.0 ± 0.2 mm/3 days for ein 2-1. Alternatively, plants may be sprayed with ethaphon or ethrel. By roots, as used here, it is

10 meant mature roots (that is, roots of any plant beyond the rudimentary root of the seedling), as well as roots and root radicles of seedlings. Stems include hypocotyls of immature plants of seedlings and stems, and plant axes of mature plants (that is, any stem beyond the hypocotyl of

15 seedlings). See Figure 7A and Figure 7B.

Ethylene sensitive wild type plants experience a shut down or diminution of endogenous ethylene production, upon application of high concentrations of ethylene. In the ethylene insensitive plants of the present invention,

20 the plants continue endogenous production of ethylene, despite administration of inhibitory amounts of ethylene. Ethylene production for wild type and ethylene insensitive mutants are shown in Table 1. An ethylene insensitive plant will produce an amount or have a rate of ethylene

25 production greater than that of a wild type plant upon administration of an inhibitory amount of ethylene. As one skilled in the art will recognize, absolute levels of ethylene produced will change with growth conditions.

Ein1 and ein2 mutants are described for example

30 in, Guzman et al., "Exploiting the Triple Response of Arabidopsis to Identify Ethylene-Related Mutants", *The Plant Cell* 1990, 2, 513, the disclosures of which are hereby incorporated herein by reference, in their entirety.

The present invention is further described in the

35 following examples. These examples are not to be construed as limiting the scope of the appended claims.

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EXAMPLE 1**PRODUCTION OF *Arabidopsis* MUTANTS**

The production of plants which exhibit enhanced disease tolerance and ethylene insensitivity were investigated with the use of *Arabidopsis* mutants *ein*, which are insensitive to ethylene and are derived from *Arabidopsis* Col-0. The *ein* mutants were prepared according to the method of Guzman et al., *The Plant Cell*, 1990, 2, 513, the disclosures of which are hereby incorporated herein by reference, in their entirety. Specifically, twenty five independent ethylene-insensitive mutants were isolated; six mutants which showed at least three-fold difference in the length of the hypocotyl compared with ethylene-treated wild-type hypocotyl, were further characterized. In these mutants, the apical hook was either present, absent or showed some curvature in the apical region. The appearance of the apical curvature was dependent on the duration of the incubation. After more than 3 days of incubation in the dark with 10 μ l/L ethylene, the apical curvature was absent. This phenotype was named "*ein*" for ethylene insensitive.

Mendelian analysis indicated that insensitivity to ethylene was inherited as either a dominant or recessive trait depending on the mutation studied. Complementation analysis was performed with five recessive mutants to determine whether more than one locus was involved in this phenotype. The results of these studies indicated that all five recessive mutations were allelic. The *ein* phenotype was tested for linkage to nine visible markers to determine whether the recessive and dominant *ein* mutations were allelic. The dominant *ein* mutation was mapped close to the mutation *ap-1* locus on chromosome 1 and was named *ein1-1*. None of the nine markers showed linkage to the recessive *ein* mutation. Restriction fragment length polymorphism (RFLP) analysis was performed to map this mutation. Randomly selected RFLP probes were initially used to assess linkage. After testing probes from three different

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chromosomes, linkage was detected to one RFLP from chromosome 4 and named ein2-1. This observation was confirmed using additional RFLP probes from the same chromosome. Further experimentation confirmed ein2-2,
5 ein2-3, ein2-4 and ein2-5 to be alleles of ein2-1.

Growth features of ethylene insensitive mutants were also observed. After seedlings were planted in soil and cold treated at 4°C for 4 days, the seedlings were incubated in the dark at 23°C for 66-72 hours. Plants were
10 grown to maturity in a growth chamber at 22°C to 25°C under continuous illumination with fluorescent and incandescent light. The rosette of ein1-1 and ein2-1 plants was larger compared with the wild type, Col-0, rosette and a delay in bolting (1 cm to 2 cm growth in the length of the stem) was
15 observed. These observations indicated that the ethylene insensitive mutations identified at the seedling stage exerted remarkable effects during adult stages of growth.

eto mutants, which constitutively produce ethylene, were initially screened by observing a
20 constitutive triple response; seedlings with inhibition of hypocotyl and root elongation, swelling of the hypocotyl and exaggerated tightening of the apical hook. Mendelian segregation analysis determined the genetic basis of these mutations to be a single recessive mutation and identified
25 as an ethylene overproducer or eto.

eto1, ein1 and ein2 mutants were analyzed to determine ethylene accumulation. The mutants were backcrossed to the wild type before physiological examination. Surface-sterilized seeds (about 500) were
30 germinated and grown for 66 to 72 hours in the dark at 23°C in 20 ml gas chromatograph vials containing 15 ml of growth medium.

To measure the conversion of exogenous 1-aminocyclopropane-1-carboxylic acid (ACC, an intermediate
35 in ethylene production) to ethylene, seedlings were grown in 1% low-melting-point agarose buffered with 3 mM Mes at pH 5.8. In this solid support no chemical formation of

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ethylene from ACC was detected at any of the concentrations of ACC employed.

Ethylene accumulation from tissues of mature plants (100 mg) was measured after overnight incubation in 20 ml gas chromatograph vials. Leaves and inflorescence were taken from 24-28 day old plants, siliques from 32-36 day old plants. Accumulation of ethylene was determined by gas chromatography using a photo-ionization detector (HNU) and a Hewlett Packard HP5890A gas chromatograph equipped with an automated headspace sampler. A certified standard of 10 μ l/L ethylene (Airco) was used to calculate ethylene concentrations. The concentration of the inhibitors of ethylene biosynthesis and ethylene action was determined empirically. For eto mutants, AVG, α -aminoisobutyric acid, and AgNO₃, supplemented the media at 5 μ M, 2mM and 0.1 mM, respectively and trans-cyclooctene (17 μ l/L) was injected into the vial after the cold treatment. Ethylene production was increased significantly in the dominant ein1-1 mutant and the recessive ein2-1 mutant, see Table 1. Ethylene production was inhibited in eto1-1 seedlings that were grown in media supplemented with ethylene inhibitors aminoethoxyvinylglycine, AGV and α -aminoisobutyric acid, AIB, see Table 1.

The EIL sequences represent cDNA sequences similar to the EIN3 sequence. They were obtained by screening an *Arabidopsis* seedling cDNA library (Kieber et al., *Cell*, 1993, 72, 427-441, at low stringency in the following manner. The cDNA library was hybridized with the radiolabeled EIN3 cDNA insert at 42° C for 48 hours in a hybridization solution consisting of 30% formamide, 5X Denhardt's solution, 0.5% SDS, 5X SSPE, 0.1 mg/ml sheared salmon sperm DNA, according to the methods of Feinberg and Vogelstein, *Anal. Biochem.* 1984, 177, 266-267, incorporated herein by reference in its entirety. The filters were washed at 42° C with 30% formamide, 0.5% SDS, 5X SSPE; followed by 2X SSPE.

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Mutageneized *HLS1* plants were obtained as set forth above for *EIN2*, *EIN3*, and *EIL*.

Table 1
Ethylene Production in Triple Response Mutants

5	Strain	Ethylene Accumulation
	Wild Type	
	Etiolated Seedlings	6.7 \pm 0.68 nL
	Light-grown Seedlings	84.25 \pm 13.95 nL
	Leaves	73.01 \pm 17.64 nL/g
10	Siliques	144.96 \pm 28.99 nL/g
	Inflorescence	234.53 \pm 18.04 nL/g
	<i>etol-1</i>	
	Etiolated Seedlings	276.72 \pm 53.70 nL
	Light-Grown Seedlings	182.01 \pm 24.84 nL
15	Leaves	174.39 \pm 29.18 nL/g
	Siliques	322.16 \pm 38.66 nL/g
	Inflorescence	1061.84 \pm 72.16 nL/g
	<i>hls1-1</i>	
	Etiolated seedlings	5.81 \pm 0.32 nL
20	Leaves	31.56 \pm 0.32 nL
	<i>ein1-1</i>	
	Etiolated Seedlings	12.73 \pm 2.79 nL
	Leaves	222.95 \pm 2.79 nL
	<i>ein2-1</i>	
25	Etiolated Seedlings	20.69 \pm 2.09 nL
	Leaves	135.59 \pm 26.89 nL/g

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Another ethylene insensitive mutant of *Arabidopsis thaliana* was designated *etr* by Bleecker et al. in "Insensitivity to Ethylene Conferred by a Dominant Mutation in *Arabidopsis thaliana*", *Science* 1990, 241, 1086, the disclosures of which are hereby incorporated herein by reference, in their entirety. *Etr* was identified by the ethylene-mediated inhibition of hypocotyl elongation in dark-grown seedlings. Populations of M_1 generation from mutagenized seed of *Arabidopsis thaliana* were plated on a minimal medium solidified with 1% agar and placed in a chamber through which 5 $\mu\text{l/L}$ ethylene in air was circulated. Seedlings that had grown more than 1 cm after 4 days were selected as potential ethylene insensitive mutants. A screen of 75,000 seedlings yielded three mutant lines that showed heritable insensitivity to ethylene. Hypocotyl elongation of *etr* mutant line was unaffected by ethylene at concentrations of up to 100 $\mu\text{l/L}$, while elongation of the wild type was inhibited by 70% with ethylene at 1 $\mu\text{l/L}$.

20 EXAMPLE 2

CLONING AND SEQUENCING OF *EIN2*

The *EIN2* locus was identified by a mapped based cloning strategy described as follows. The *ein2-1* mutant was crossed onto the DP28 marker line (*dis1*, *clv2*, *er*, *tt5*) according to the methods of Koornneef and Stamm, *Methods in Arabidopsis Research*, eds. C. Koncz, N-H Chua, and J. Schell, 1992, World Scientific Publishing Co., Singapore, incorporated herein by reference in its entirety. The F_2 progeny were mapped with Restriction Fragment Length Polymorphisms (RFLPs) according to the methods of Chang et al., *Proc. Natl Acad. Sci. USA* 1988, 85, 6856 and Nam et al., *Plant Cell* 1990, 1, 699, the disclosures of which are hereby incorporated by reference in their entirety.

The *ein2-1* mutation was found to segregate with RFLPs on the top of chromosome five (Table 2). Two recombinant progeny found with $\lambda 217$ (E15 and E54) were also

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recombinant with the more proximal g3837 and λ 291 clones, indicating that *ein2-1* is distal to λ 217. Recombinant plants were identified by examining F_1 families from the *ein2-1* x DP28 cross for the genotype at the λ 217 locus.

5 Protocols are the same mapping with RFLPs. Recombinants were defined by having at least one recombinant chromosome in an *ein2-1* homozygote. The Ubq6121 marker, however, identified a different F_2 progeny (E46) as being recombinant. This positions *ein2* within the interval of

10 λ 217 and Ubq6121. To further limit the position of *ein2* on the top of chromosome 5, recombinants were sought with the PCR based marker ATHCTR1, Bell et al., *Methods in Plant Molecular Biology: A Laboratory Manual*, 1993, eds. Maliga, Klessig, and Cashmore, Cold Spring Harbor Laboratory Press,

15 the disclosure of which is hereby incorporated by reference in its entirety.

A single recombinant progeny was identified in 102 F_2 progeny scored. This F_2 progeny was also recombinant at the proximal λ 217 and ASA1 markers,

20 demonstrating the position of *ein2* as distal to ATHCTR1. Additional genetic information was generated by examining recombinant progeny from a cross between *ein2-1* and *hy5*. Two additional recombination events between *ein2-1* and ATHCTR1 were identified by this approach. There were no

25 recombinant plants identified at the g3715 locus, a cosmid clone identified in Nam et al., *supra*.

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Table 2
Characterization of Plants Having ein2 Mutation

	ALLELE	HYPOCOTYL	SE	ROOT	SE	TL	SE
	Columbia	3.6	0.2	1.6	0.1	5.2	0.2
5	Landsberg	3.2	0.1	1.7	0.1	4.9	0.2
	Wassilewskija	2.7	0.1	0.9	0.1	3.6	0.1
	ein2-1 *	6.0	0.3	7.1	0.1	13.1	0.4
	ein2-3 *	8.2	0.2	5.9	0.3	14.1	0.4
	ein2-4 *	7.5	0.2	6.3	0.4	13.8	0.5
10	ein2-5 *	8.4	0.2	7.2	0.5	15.6	0.5
	ein2-6	8.8	0.4	5.4	0.2	14.2	0.5
	ein2-7	5.9	0.1	3.8	0.1	9.7	0.2
	ein2-9	7.3	0.2	5.5	0.2	12.8	0.3
	ein2-10	6.4	0.1	4.7	0.4	11.1	0.5
15	ein2-11	8.1	0.1	7.7	0.3	15.8	0.4
	ein2-12	6.5	0.3	4.4	0.3	10.9	0.4
	ein2-13	5.4	0.2	3.7	0.2	9.1	0.4
	ein2-15	6.9	0.5	5.3	0.4	12.2	0.9
	ein2-16	8.1	0.3	7.7	0.6	15.8	0.7
20	ein2-18 +	6.2	0.2	6.5	0.4	12.7	0.4
	ein2-19 +	7.1	0.2	6.2	0.5	13.3	0.6
	ein2-20 +	5.8	0.2	5.2	0.2	11.0	0.3

All units are in mm, TL = Total Length, SE = Standard Error
 * Guzman and Ecker, Plant Cell 1990, 2, 513.

25 + Gift of Caren Chang and Elliot Meyerowitz, Pasadena, CA.

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The flanking genetic markers were used to build a Yeast Artificial Chromosome (YAC) physical contig spanning the *ein2* locus (Figure 1). The YAC positions were identified by colony hybridization pursuant to the

5 technique of Matallana, et al., *Methods in Arabidopsis Research*, eds C. Koncz, N-H Chua, and J Schell, 1992, World Scientific Publishing Co., Singapore, the disclosures of which are hereby incorporated by reference in their entirety.

10 YAC clones are replicated in the yeast cells as authentic chromosomes and so they are present as only one copy per cell. This is an important difference with bacterial colony hybridization and makes colony filter treatment a critical step for successful sequence

15 detection. After growing colonies overnight on the filters, the cell walls were digested and the spheroplasts were lysed in order to prepare yeast DNA for hybridization.

Yeast cell wall digestion is stimulated by reducing agents, such as 2-mercaptoethanol or DTT, that

20 modify the wall structure and make it more sensitive to enzymatic action. Colony filters were placed on filter paper soaked in 0.8% DTT in SOE buffer (1 M sorbitol, 20 mM EDTA, 10 mM Tris-acetate pH 8.0) for 2-3 min. before transferring them to filter paper soaked in SOE containing

25 1% 2-mercaptoethanol and 1 mg/ml Zymolyase 10-T in individual 150 X 15 mm petri dishes. Petri dishes were parafilmed and stacked in a sealed plastic bag and incubated at 37° C overnight.

After spheroplasting, lysis was carried out by

30 placing the filters on whole sheets of Whatman 3MM paper soaked in the appropriate solution. The 3MM sheets were placed on Saran wrap and soaked immediately before use. The filters were treated as follows:

1. 10% SDS for 10 min.;
- 35 2. 0.5 M NaOH for 10 min (1.5 NaCl should be included for Hybond N+); Repeat;
3. Air dry for 5 min.;

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4. 1 M Tris-HCl (pH 7.6), 1.5 M NaCl for at least 5 min;

5. 0.1 M Tris-HCl (pH 7.6), 0.15 M NaCl for at least 5 min. Cell debris on the filters was eliminated by gently wiping the filters with Kimwipes soaked in the same solution.

6. 2xSSPE for at least 5 min. This step precedes hybridization. Following lysis, the filters are air dried for 30 min. and baked for 2 hours at 80 C.

10 The left ends of the identified YAC clones were isolated by plasmid rescue according to Bell et al., 1994. Right ends were isolated by either vectorette PCR according to the methods of Matallana, et al., 1992, *supra*. or inverse PCR as described by Bell, et al., 1994, *supra*, the
15 disclosures of which are hereby incorporated by reference in their entirety. The yUP library appeared to be missing clones corresponding to ATHCTR1; three clones hybridizing to this locus were found within the EG library (Grill and Somerville, Mol. Gen. Genet. 1991, 226, 484, incorporated
20 herein by reference in its entirety.) The pEG23G5L left end plasmid rescue hybridizes to useful EcoR I and Xba I polymorphisms and hybridizes to the same lambda clone as ATHCTR1 (λ ctg24; Kieber et al., Cell 1993, 72, 427, incorporated herein by reference in its entirety). The
25 left end rescue pyUP2G11L hybridizes to EG23G5, linking the Ubq6121/g3715 and ATHCTR1 clones into a contiguous array. pyUP2G11L also contains a Bgl II polymorphism that is informative in the ein2-1 X DP28 cross. The three plants that are recombinant at ATHCTR1 are also recombinant at
30 pyUP2G11L; this indicates the position of ein2 is distal to this YAC end (Figure 1).

To facilitate the identification of the ein2 locus, 24 alleles were identified (Table 1; Guzman and Ecker, Plant Cell 1990, 2, 513, incorporated herein by
35 reference in its entirety.) Many of these alleles were generated by X-ray or diepoxybutane mutagenesis; these mutagens are known to create polymorphisms that are

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detectable by hybridization to a genomic Southern blot (Clark, et al., *Genetics* 1986, 112, 755; Reardon et al., *Genetics* 1987, 115, 323, incorporated herein by reference in their entirety). *EcoR I*, *Hind III*, *BamH I*, *Bgl II*, and *Sal I* genomic Southern blots were made to find such a polymorphism in the mutant alleles of *ein2*. The following probes that mapped between Ubq6121 and yUP2G11L were hybridized to the genomic allele blots: Ubq6121, EG19A10L, yUP2G11R, g3715, yUP19E11L, EG23G5R, and yUP2G11L. The cosmid clone g3715 hybridized to a restriction fragment length polymorphism in *ein2-12* that corresponds to a lost *EcoR I* site (Figure 2). Based on this missing *EcoR I* site, this region was examined further.

The 1.2 kb *EcoR I* fragment that corresponds to one of the missing bands in *ein2-12* was subcloned from g3715 into pKS (Stratagene, LaJolla, CA) this clone is named pgEE1.2 (Figure 3). The pgEE1.2 insert was used to isolate 22 cDNA clones made from ethylene treated three-day old etiolated *Arabidopsis thaliana* seedlings (Kieber, et al. 1993, *supra*.) pgEE1.2 was also used to identify a single genomic lambda clone, λ gE2, from a λ DASH II library made from adult Columbia plants. The λ gE2 clone spanned the 5' end of the locus and terminated within the 3' end of the cDNA. Initially the pCE2.5 clone was sequenced but since this clone was not full length, the 5' ends of pCE2.17, pCE2.20, and pCE2.22 (Kieber, et al. 1993) were sequenced to determine the structure of the full length frame and ending within 60 bp from a putative "TATA" box (Figure 4). Using 5 μ g of poly(A+) RNA from 3-day old dark-grown, ethylene-treated *Arabidopsis* seedlings (hypocotyls and cotyledons) as template and oligo(dT) as primer, first-strand cDNA synthesis was catalyzed by Moloney murine leukemia virus reverse transcriptase (Pharmacia) for construction of the *Arabidopsis* cDNA expression library. Second-strand cDNA was made as described by Gubler and Hoffman, *Gene* 1983, 25, 263, which is hereby incorporated by reference in its entirety, except

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that *E. coli* DNA ligase was omitted. After the second-strand reaction, the ends of the cDNA were made blunt with Klenow fragment, and *EcoR* I-Not I adaptors (Pharmacia) were ligated to each end. The cDNA was purified from unligated
5 adaptors by spun-column chromatography using Sephacryl S-300 and size fractionated on a 1% low melting point minigel. Size-selected cDNAs (0.5-1, 1-2, 2-3, and 3-6 kb) were removed from the gel using agarose (New England BioLabs), phenol-chloroform extracted, and precipitated
10 using 0.3M NaOAc (pH 7)-ethanol. A portion of each cDNA size fraction (0.1 µg) was coprecipitated with 1 µg of λZAPII *EcoR* I-digested, dephosphorylated arms and then ligated overnight in a volume of 4 µl. Each ligation mix
15 was packaged in vitro using Gigapack II Gold packaging extract (Stratagene). The structure of this locus was determined by Southern hybridization and restriction mapping of the λgE2 and g3715.

The sequence of the *EIN2* genomic DNA was determined from PCR products and the λgE2 genomic lambda
20 clone. Primers were selected from the sequence of the pcE2.5, pcE2.17, and genomic subclones of λgE2. The primers were then commercially synthesized (Research Genetics, Huntsville, AL).

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Table 3
PRIMERS FOR THE E1N2 LOCUS

	SEQUENCE ID NO.	Primer Name	Sequence	position
5	21	PE2.7A	GGATCCTCTAGTCAAATTACCGC	
	22	PE2.7B	AGATCTGGTATATTCCGTCTGCAC	
	23	PE2.5'	CCGGATTTCGGTTTGTAGC	PCR/ 3' end
	24	PE1	GACGTGCATGTTCTTGGG	
	25	PE2	GAAAGCCACATCACCTGC	
10	26	PE3	GGGGTGGAGTTATCCAC	
	27	PE4	GACACCGGGAAGTATCG	
	28	PE5	CTGCTTTCATAGAAGAGGC	PCR/ middle
	29	PE6	GTCAGAACAAACCTGCTCC	PCR/ 5' end
	30	PE7	CACCCAGGTCTTGGTGG	
15	31	PE8	GGCCGCCATGGATGCG	
	32	PE9	TCTCAATCAAGAGGAGGC	
	33	PE10A	CTTGAAGGATCCGAGTGG	
	34	PE11	CAGGTTGGCGAGTTCCTCG	
	35	PE12	CTTGCTGTTATTCTCCATGC	
20	36	PE13	CCCTGGACCAGCTCCTGG	
	37	PE14	TGGCGCAAGCATCGTCCC	PCR/ middle
	38	PE15	AAATGTTTCAGGAATCTCTCG	
	39	PE16	CTGGCTGGCAGCCACGCC	PCR/ 3' end

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	40	PE17	GCGTTCTCAAAGCTGCGG	
	41	PE18	ACTGATGGGTCTTCTGGG	
	42	PE19	GGATCAGGATGGACCCGG	
	43	PE20	TGGTTGCTGAAGCCAGGG	
5	44	PE21	TCCATT CATAGAGAGTGGG	
	45	PE22	ATGCCCAAGAACATGCACG	
	46	PE23	CAACTGATCCTTTACCCTGC	
	47	PE24	GTTGTTAGGTCAACTTGCG	PCR/ 5' end
	48	PE25	CTCTGTTAGGGCTTCCTCC	
10	49	PE26A	GAATCAGATTTCGCGAGG	
	50	PE27	GTCCAAATGGAGGAAGCC	
	51	PE28	CCACGACTGTACAATTGACCTTG	engine- ered MunI site
	52	PE29	CATGATCGCAAGTTGACC	
	53	PE30	AGAAACTCTTATCAAGCTACG	
15	54	PE31	AAGCTTATGGGTGCTCGTGC	
	55	PE32	GGAAAGAGAGAAAGACTCAG	
	56	PE33	GCCACCAAGTCATACCCG	

Primer sequences are set forth 5' to 3'.

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Four overlapping regions of the *ein2* locus between 1.2 and 3.2 kb in length were rapidly amplified by polymerase chain reactions (Idaho Technologies, Idaho falls, Idaho). Conditions for the PCR reactions are as follows: 92°C, 2 seconds; 56°C, 2 seconds; 72°C, 1 minute; 50 cycles. Between 200 and 500 ng of these PCR products were directly sequenced on the ABI373A automated sequencer using Taq Dye-Terminator chemistry (Applied Biosystems Division, PEC). The genomic sequence of the wild type Columbia *EIN2* locus is shown in Figure 5. Eight mutant alleles of *ein2* were also sequenced and the corresponding mutations identified (Table 4). The presence of these mutations in the mutant alleles of *ein2* confirms the identity of this gene as *EIN2*.

Table 4
IDENTIFIED MUTATIONS OF *EIN-2*

ALLELE	MUTAGEN	MUTATION	POSITION*	RESULT
<i>ein2-3</i>	X-ray	Insert T	+3642	Frameshift
<i>ein2-4</i>	X-ray	AG to TT	+2103	Frameshift
<i>ein2-5</i>	X-ray	ΔCATGACT	+1570	Frameshift
<i>ein2-6</i>	Agro-bacterium	ΔGAGTTGCGC ATG (SEQ ID NO: 17)	+965	ΔGVAH (115) (SEQ ID NO: 18)
<i>ein2-9</i>	DEB	A to C	+4048	H to P
<i>ein2-11</i>	DEB	TG to AT	+3492	Ochre
<i>ein2-12</i>	X-ray	ΔTGCTACAAT CAGAATTCTT GCAGT (SEQ ID NO: 19)	+1611	ΔATIRILAV (SEQ ID NO: 20)
<i>ein2-16</i>	X-ray	AGT to G	+2851	Frameshift

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* Position relative to the start of pCE2.17; see Figure 5, nucleic acid; position 1 corresponds to the beginning of the cDNA.

EXAMPLE 3

5 CLONING AND SEQUENCING OF EIN3

In order to clone the EIN3 gene a collection of 5000 T-DNA insertion lines (Feldmann and Marks, *Mol. Gen. Genet.* 1987, 208, 1-9, incorporated herein by reference in its entirety) was screened for ethylene-insensitive mutants. A mutant with a phenotype similar to that of ein3-1 (an EMS generated allele) was identified and genetic complementation tests revealed that ein3-1 and the T-DNA insertion mutant (designated ein3-2) were allelic. Complete cosegregation of the mutant phenotype and the dominant kanamycin resistance marker on the T-DNA indicated that the T-DNA insertion was located within, or at least very close, to the EIN3 gene. Genomic DNA flanking the T-DNA insert was cloned using the left border rescue technique. Genomic Southern blots of wild-type and ein3-2 DNA hybridized with the rescued fragment indicated that the cloned segment of Arabidopsis DNA corresponded to sequences disrupted by the T-DNA insert and did not result from cloning an unlinked fragment of genomic DNA. In all restriction digests the mobility of the hybridizing fragments is shifted in the insertion mutant relative to wild-type.

cDNA and genomic libraries constructed from wild-type DNA were screened with the rescued DNA fragment. The cDNAs obtained indicated that the EIN3 gene encodes a 628 amino acid open reading frame. Structural features of the predicted polypeptide include: 1) a region rich in acidic amino acids at the amino terminus, 2) several basic domains in the central portion of the protein, and 3) several poly-asparagine repeats near the carboxy terminus. Although database searches revealed no overall similarities to any characterized proteins, the three structural motifs described are found in transcriptional regulatory proteins.

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Stretches of acidic amino acids function in transcriptional activation presumably through binding to other proteins. Basic domains serve as nuclear localization signals and can bind DNA. Poly asparagine repeats are present in the SWI1 protein of yeast. This protein has been termed a transcriptional accessory protein because it is required for transcriptional activation of target genes but does not bind directly to DNA. It has been suggested that the poly asparagine repeats are involved in protein-protein interactions.

Sequencing genomic clones indicated that the EIN3 gene has a very simple structure. There are no introns within its open reading frame. However there is a single intron located in the 5' transcribed region. In addition to sequencing the wild-type EIN3 gene, genes from three independently isolated ein3 mutants were sequenced. In each case an alteration was identified confirming the identification of the bona fide EIN3 gene. In the ein3-1 allele, a point mutation introduces a premature in frame stop codon. The ein3-2 allele contains a T-DNA insertion which interrupts the coding region. A point mutation in the ein3-3 allele substitutes an acidic amino acid for a basic amino acid within one of the basic regions described above.

The expression pattern of the EIN3 gene in seedlings was examined by placing the GUS reporter gene under control of the EIN3 promoter. The construct employed was a translational fusion including 5' non-transcribed sequences, the 5' intron and 93 amino acids of the EIN3 coding region cloned upstream of the GUS gene in the pBI101 vector (Jefferson et al., *EMBO J*, 1987, 6, 3901-3907, incorporated herein by reference in its entirety) and named pHSEIN3GUS. *Arabidopsis* root explants were transformed and transgenic plants regenerated (Velvickins et al., *PNAS* 1988, 85, 5536-5540, incorporated herein by reference in its entirety). The GUS activity patterns observed suggest that the EIN3 promoter is most active in expanding or elongating cells. In three day old etiolated seedlings GUS activity

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staining is located predominantly in the apical hook and root tips. In younger seedlings in which the hypocotyl is not fully extended staining is also prevalent throughout this tissue. In 14 day old light grown seedlings abundant
5 GUS activity is observed in the roots, upper portions of the hypocotyl, cotyledons and leaves. The EIN3 promoter is not induced by ethylene as the levels of GUS activity in air and ethylene treated seedlings appear equivalent. This observation is supported by the fact that steady state
10 levels of the endogenous EIN3 transcript are similar in ethylene and air treated seedlings and adult plants as determined by Northern analysis.

The EIN3 coding region was cloned downstream of the bacterial reporter gene B glucuronidase (GUS) in the
15 plasmid pRTL2-GUS according to the methods of Restrepo et al., *Plant Cell* 1990, 2, 987-998, incorporated herein by reference in its entirety, to create pNLEIN3Bgl2 (see Figure ____). The plasmid was transformed into *Arabidopsis* protoplasts and transiently expressed according to the
20 methods of Abel and Theologis, *Plant J.* 1994, 5, 421-427, incorporated herein by reference in its entirety. All detectable GUS activity was targeted to the nuclei of the protoplasts indicating that the EIN3 protein functions in the nucleus. These results suggest that the EIN3 protein
25 may function as a transcription factor which regulates ethylene-regulated gene expression.

The EIN3 gene is a member of a small gene family. Low stringency hybridization of genomic Southern blots indicates that there are at least two members in addition
30 to EIN3. Three EIN3 homologue, designated as EIL1, EIL2, and EIL3, have been cloned and sequenced. The EIL and EIN3 predicted polypeptides structurally similar in that the amino termini of both proteins are rich in acidic amino acids and their central regions contain several basic
35 domains. Their carboxyl termini are not as well conserved as EIL1 contains a polyglutamine repeat instead of poly asparagine repeats. The EIL2 and EIL3 polypeptides do not

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contain polyglutamine repeats or poly asparagine repeats. It is interesting to note that the amino acid substitution in the ein3-3 allele occurs in one of the regions rich in basic amino acids that is completely conserved between the EIN3 and EIL polypeptides. Currently, it is not known whether the EIL gene product functions in the ethylene signal transduction pathway of Arabidopsis. However at this time, the EIL1 and EIL2 cDNAs do not map to the same location as any of the characterized ethylene response mutations. The location of the EIL3 cDNA has not yet been mapped. The EIL1 polypeptide is the most similar to EIN3.

The ein3 mutant alleles were sequenced on an Applied Biosystems 373A DNA Sequencing System (Foster City, CA) using Tag dideoxy terminator chemistry (Applied Biosystems). The PCR primers are set forth in Table 5.

TABLE 5
PRIMERS FOR EIN3 PCR

SEQUENCE ID NO.	PRIMER NAME	SEQUENCE	POSITION in genomic
57	PR24	CCTTCTATATTTGGTTCC	680-698
58	PR15	CCATTCTCCGGAATAATCC	1306-1324
59	PR5	CACGGAGCAGGATAAGGGTA	1148-1166
60	PR19	CGGATTGGATTGTGTGTGC	3312-3331

The primer sequences are set forth 5' to 3'.

Primer pairs PR24 - PR15 and PR5 - PR19 were used to amplify genomic DNA from the ein3 mutants. PCR amplification was performed with a Bioscycler Oven (New Haven, CT). Conditions for amplification were as follows: 92° C for 1 min; 55° C for 1 min.; 72° C for 3 min. The mutations discovered are listed in Table 6.

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Table 6

IDENTIFIED MUTATIONS OF EIN3

Allele	Mutagen	Sequence change	Consequences of sequence change
ein3-1	EMS	G to A, position 1598	amino acid 215, W to umber
5 ein3-2	T-DNA	position 2001	T-DNA insertion
ein3-3	DEB	G to T, position 1688	amino acid 245, K to N

The EIN genes were obtained by screening an *Arabidopsis* seedling cDNA library (Kieber et al., Cell, 1993, 72, 427-441, at low stringency in the following manner. The cDNA library was hybridized with the radiolabeled EIN3 cDNA insert at 42° C for 48 hours in a hybridization solution consisting of 30% formamide, 5X Denhardt's solution, 0.5% SDS, 5X SSPE, 0.1 mg/ml sheared salmon sperm DNA, according to the methods of Feinberg and Vogelstein, Anal. Biochem. 1984, 177, 266-267, incorporated herein by reference in its entirety. The filters were washed at 42° C with 30% formamide, 0.55 SDS (should this be 0.5% SDS?), 5X SSPE; followed by 2X SSPE.

EXAMPLE 4

20 HOOKLESS MUTATION OF THE APICAL HOOK

The "triple response" in *Arabidopsis thaliana* occurs in response to the plant hormone ethylene and is characterized by three distinct changes in the morphology of etiolated seedlings. These include, exaggeration of the apical hook, radial swelling of the hypocotyl, and inhibition of root and hypocotyl elongation. Observation

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of the apical hook was recorded by Charles Darwin as early as 1896.

The hook causes the apical portion of the seedling to become nearly parallel with the basal portion.

5 Production of the bend in the hypocotyl requires either a larger number of cells, or increased elongation of cells on the adaxial side (outside) of the hook. A study of the characteristics of hook formation in bean seedlings demonstrated that the curvature is produced by differential

10 growth rates on each half of the hypocotyl resulting in longer cells on the convex side of the hook, see Rubenstein, 1972 *Plant Physiology* 49:640-643.

Previous studies suggest that hormones may be involved in hook formation. The hormones involved are

15 believed to be auxin and ethylene. Auxin is known to be a controlling factor in cell elongation in the hypocotyl, see Klee and Estelle, 1991 *Annual Review of Plant Physiology* 42:529-551, incorporated herein by reference in its entirety, and ethylene has been shown to exaggerate the

20 bending of the hook in wild type etiolated seedlings (Guzman and Ecker, *supra*). One hypothesis to explain hook formation is that auxin promotes elongation of cells on the outside of the apical hook allowing differential growth rates and bending. Work performed by McClure and Guifoyle

25 (1989) demonstrated that the initial uniform expression of small auxin up-RNA (SAUR) mRNA on both sides of the hypocotyl was altered when the tissue was transferred from an erect to horizontal position. An increase in SAUR mRNA accumulation was observed on the "outside" region and a

30 concurrent rapid decrease in SAUR mRNA occurred on the "inside" region of an upward bending hypocotyl. Ethylene has been shown to alter transport of auxin in hypocotyl tissue (Mattoo and Suttle, *supra*), suggesting a possible role for ethylene in exaggeration of the hook. To

35 exaggerate the hook, ethylene might affect auxin localization causing even more bending on the outside of the hook.

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The triple response of *Arabidopsis* has been used to isolate mutants affected in the ethylene response. The *hookless 1(hls1)* mutant exhibits a tissue specific defect in the triple response. Null mutants (*hls1-1*) completely
5 lack the apical hook in the presence and absence of ethylene while weak alleles of *hls1* (*hls1-2*) show some bending in the hook in the presence of ethylene. The complementation cross between *hls1-1* and *hls1-2* gave rise to F1 progeny which resembled *hls1-2*. In addition to *hls1-1*
10 and *hls1-2*, six EMS alleles, three DEB alleles, one X-ray allele, and two non-tagged T-DNA alleles have been isolated in accordance with the methods set forth in Guzman et al. The *Plant Cell* 1990 2:513-523, hereby incorporated by reference in its entirety (Table 7). Seven of these are
15 strong alleles which are completely hookless in the presence of ethylene. Five of these are weak alleles showing a partial bend in the presence of ethylene. The *hls1* phenotype is epistatic in the hook with other ethylene mutants.

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Table 7
IDENTIFIED PHENOTYPIC AND PROTEIN MUTATIONS OF HLS1

	ALLELE	MUTAGEN	HOOK ANGLE	CHANGE
	<i>hls1-1</i>	EMS	2.2 ± 0.9	aa345 E to K
5	<i>hls1-2</i>	T-DNA	26.2 ± 3.2	T-DNA insertion
	<i>hls1-3</i>	X-RAY	8.1 ± 1.8	4.8kb deletion of promoter
	<i>hls1-4</i>	DEB	ND (strong)	aa345 E to K
	<i>hls1-5</i>	DEB	1.3 ± 0.5	splice donor site mutated
	<i>hls1-6</i>	EMS	2.1 ± 1.0	aa326 K to W
10	<i>hls1-7</i>	DEB	3.0 ± 1.3	splice donor site mutated
	<i>hls1-8</i>	EMS	2.1 ± 1.2	aa180 R to stop
	<i>hls1-9</i>	EMS	6.3 ± 1.5	aa11 R to stop
	<i>hls1-10</i>	EMS	23.2 ± 3.0	aa1 M to I
	<i>hls1-11</i>	T-DNA	3.0 ± 1.2	ND
15	<i>hls1-12</i>	EMS	ND (weak)	NC
	<i>hls1-13</i>	EMS	ND (weak)	NC
	<i>hls1-14</i>	T-DNA	ND (strong)	ND

ND = not determined;

NC = no change in coding region or introns

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Gene Structure and Analysis

The *HLS1* gene was cloned by left border rescue of a T-DNA inserted in the promoter of *hls1-2*. The rescued fragment was used to isolate a 12kb genomic clone which was
5 then used to isolate three cDNA clones. The T-DNA was found to have inserted 710bp upstream from the 5' end of a 1.7kb cDNA clone. Deletions of the 1.7kb cDNA clone were generated in both directions using Exonuclease III. These clones were sequenced using Sequenase 2.0. Deletions of
10 the genomic clone were also generated using Exonuclease III. These clones were also sequenced. The sequence of the genomic clone covered the entire 1.7kb cDNA as well as 1712bp upstream of the start of the cDNA and 313 bp at the 3' end of the cDNA. This gene has two introns of 342 bp
15 and 81bp in size. The cDNA encoded a 403 amino acid protein of about 43kDa.

Sequence Analysis of the Alleles

The *hls1* gene from ten of the fourteen alleles was sequenced. The transcribed region as well as both
20 introns were sequenced. The *hls1* gene from each allele was isolated by PCR amplification. The sequences of the primers is set forth in Table 8.

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Table 8
PRIMERS FOR HLS1 PCR

SEQUENCE ID NO.	PRIMER NAME	SEQUENCE	POSITION in genomic
5 61	II.1	cgccactgcatgtaagaac	1303-1321
62	II.2	tccacacgcttaatacggc	3229-3211
63	II.6	ggtacggagaagaaggag	2546-2563
64	III.1	cgcgggatattgattcggt	3071-3090
65	III.2	gtgttgaaacacgcccacaa	ND
10 66	III.3	acgacaccacaaccacct	3479-3462
67	III.5	gacaagaagacacaaacc	3880-3863
68	pr1	gaatcggaggagaaggtc	3386-3403

Primer sequences are set forth 5' to 3'.

- PCR was performed on a Biosycler (New Haven, CT).
- 15 Conditions were 92° C, 1 min.; 55° C, 1 min.; 72° C, 3 min. for 35 cycles. Some of the PCR products were subcloned and sequenced using Sequenase. Additional PCR products were sequenced directly using sequence specific primers and Tag sequencing on an ABI automated sequencer (Foster City, CA).
- 20 Alleles found to contain a sequence change from wild type were confirmed by direct sequencing of the PCR product along with a wild type control. The changes found in these alleles are listed below in Table 9.

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Table 9

IDENTIFIED GENOTYPIC AND PROTEIN MUTATIONS OF HLS1

ALLELE	MUTAGEN	SEQUENCE CHANGE	CONSEQUENCES OF SEQUENCE CHANGE
<i>hls1-1</i>	EMS	G to A position 3487	aa345 E to K
5 <i>hls1-5</i>	DEB	T to A position 2194	splice donor site mutated
<i>hls1-7</i>	DEB	T to A position 2194	splice donor site mutated
<i>hls1-6</i>	EMS	T to G position 3431	aa326 K to W
<i>hls1-4</i>	DEB	G to A position 3487	aa345 E to K
<i>hls1-9</i>	EMS	C to T position 2060	aa11 R to stop (CGA - TGA)
10 <i>hls1-8</i>	EMS	C to T position 2992	aa180 R to stop (CGA - TGA)
<i>hls1-10</i>	EMS	G to A position 2033	aa1 M(start) to I

Two alleles which showed no changes in the transcribed region or in the introns, *hls1-12* and *hls1-13*, were both weak alleles. *hls1-12* was found to have reduced levels of transcript compared with wild type. It is possible that there are sequence changes in the promoter region of *hls1-12* and *hls1-13*.

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Spatial and Temporal Detection and Expression

Northern analysis of the alleles revealed weak alleles *hls1-2*, *hls1-3*, *hls1-12* all show a reduction in the amount of transcript. The *HLS1* transcript was found to be up regulated by ethylene.

HLS1 Homology

Sequence comparison was done at the DNA as well as the amino acid level using Blast and TFASTA (GCG). Some homology to one class of acetyl transferases was found.

10 There are several classes of acetyl transferases with little homology between classes. The homology in one class of acetyl transferases is comprised of only a loose consensus. *HLS1* is similar to a class of acetyl transferases found in bacteria and yeast and not similar to

15 the class found in mammalian systems. Tercero, J.C., *JBC* 1992, 267, 20270, published a minimum consensus for one class of acetyl transferases. Other members of this class include yeast *MAK3* gene, which acetylates a viral coat protein and perhaps some mitochondrial proteins. The *rimL*

20 and *rimJ* proteins are also in this class of acetyl transferases. These are *E. coli* proteins which acetylate ribosomal proteins L12 and L5. Also included in this class is the *ARD1* protein of yeast. Mutants in this gene show a specific mating defect, an inability to sporulate, and loss

25 of viability in stationary phase. There are several other bacterial members of this class. The other 150 amino acids of the *HLS1* gene show no significant homology to any proteins in the database.

Various modifications of the invention in

30 addition to those shown and described herein will be apparent to those skilled in the art from the foregoing description. Such modifications are also intended to fall within the scope of the appended claims.

SEQUENCE LISTING

(1) GENERAL INFORMATION:

- (i) APPLICANT: Trustees of The University of Pennsylvania
- (ii) TITLE OF INVENTION: Plant Genes for Sensitivity to Ethylene and Pathogens
- (iii) NUMBER OF SEQUENCES: 82
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: Woodcock, Washburn, Kurtz, Mackiewicz & Norris
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 - (E) COUNTRY: USA
 - (F) ZIP: 19103
- (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.25
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER: PCT/US95/07744
 - (B) FILING DATE: 15-JUNE-1995
 - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: 08/261,822
 - (B) FILING DATE: June 17, 1994
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: Beardell, Lori Y.
 - (B) REGISTRATION NUMBER: 34,293
- (ix) TELECOMMUNICATION INFORMATION:
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 - (B) TELEFAX: (215) 568-3439

(2) INFORMATION FOR SEQ ID NO:1:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 6042 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

TTCTCTCTCT CTCTTTGAAG GTGGCACGAG CACCCATAAC CTTGAGACCT ATAGATACAA	60
ATATGTATGT ATACGTTTTT TATATATAAA TATTTTATAT AATTGATTTT TCGATCTTCT	120
TTTATCTCTC TCTTCGATG GAACTGAGCT CTTTCTCTCT TTCCTCTTCT TTTCTCTCTC	180

TATCTCTATC	TCTCGTAGCT	TGATAAGAGT	TTCTCTCTTT	TGAAGATCCG	TTTCTCTCTC	240
TCTCACTGAG	ACTATTGTTG	TTAGGTCAAC	TTGCGATCAT	GGCGATTTTCG	AAGGTGACTT	300
CTTTCAAAAA	CCCTAATCCT	CTGTTTTTTT	TTTTATTTTG	CTGGGGGGCT	TTGTACGGAC	360
TTTCATGGGT	TTTTGTAGCT	TTCCCTCGG	CTTTTGCGCA	AATGAGACTT	TCTGGGTTTT	420
TTTTCCAGCT	TTTATAAATT	TCATCAGGTG	GATCGAATTC	GTAGTTTCAG	CTTAGATCTC	480
TCTCCCTCTT	CATTATCTGG	ACTTTCCAGA	CTTGGAGTTC	TCGGGATTG	TTTTCGGTTT	540
CTGGGTTTTG	TTTAAATTGC	GAGATTTAAG	CTTTTTTCTT	TTTTACTACT	GTA CTGGTT	600
TGTGGTTGAC	CTTTTTTTTC	CTTGAAGATC	TGAATGCGTA	GATCATACGG	GATCTTTGCA	660
TTTTTGTTGC	TTTTCGTCAG	CGTTACGATT	CTTTTAGCTT	CAGTTTAGTT	GAAATTTGTA	720
TTTTTTTTGA	GCTTATCTTC	TTTTTGTTGC	TGCTTCATAC	TAAGATCAAT	TATTGATTTG	780
TAATACTACT	GTATCTGAAG	ATTTTCACCA	TAAAAAATAA	ATTCAGGTCT	GAAGCTGATT	840
TCGAATGGTT	TGGAGATATC	CGTAGTGGTT	AAGCATATGG	AAGTCTATGT	TCTGCTCTTG	900
GTTGCTCTGT	TAGGGCTTCC	TCCATTTGGA	CCAACCTAGC	TGAATGTTGT	ATGATCTCTC	960
TCCTGAAGC	AGCAAATAAG	AAGAAGGTCT	GGTCCTTAAC	TTAACATCTG	GTTACTAGAG	1020
GAAACTTCAG	CTATTATTAG	GTAAGAAAG	ACTGTACAGA	GTGTATAAC	AAGTAAGCGT	1080
TAGAGTGGCT	TTGTTTGCCT	CGGTGATAGA	AGAACCGACT	GATTCGTTGT	TGTGTGTTAG	1140
CTTTGGAGGG	AATCAGATTT	CGCGAGGGAA	GGTGTTTTAG	ATCAAATCTG	TGAATTTTAC	1200
TCAACTGAGG	CTTTTAGTGA	ACCACGACTG	TAGAGTTGAC	CTTGAATCCT	ACTCTGAGTA	1260
ATTATATTAT	CAGATAGATT	TAGGATGGAA	GCTGAAATTG	TGAATGTGAG	ACCTCAGCTA	1320
GGGTTTATCC	AGAGAATGGT	TCCTGCTCTA	CTTCCTGTCC	TTTTGGTTTC	TGTCGGATAT	1380
ATTGATCCCG	GGAAATGGGT	TGCAAATATC	GAAGGAGGTG	CTCGTTTCGG	GTATGACTTG	1440
GTGGCAATTA	CTCTGCTTTT	CAATTTTGCC	GCCATCTTAT	GCCAATATGT	TGCAGCTCGC	1500
ATAAGCGTTG	TGACTGGTAA	ACACTTGGCT	CAGGTAAACA	TTTTTCTGAT	CTCTAAAGAG	1560
CAAACTTTTT	AAAATAACAA	ACTGGGCTCT	GTGGTTGTCT	TGTCACTTTC	TCAAAGTGGA	1620
ATTCTACTAA	CCACCTTCTC	TATTTTTCTA	ACATTTTAAT	GTTCTTTACT	GGGACAGATC	1680
TGCAATGAAG	AATATGACAA	GTGGACGTGC	ATGTTCTTGG	GCATTGAGGC	GGAGTTCTCA	1740
GCAATTCTGC	TCGACCTTAC	CATGGTAGTT	ACTTACAATT	CTTGCTGTT	CTTAATTTTT	1800
TTATTATGTA	GTAAAATTTT	GATTCCTCTG	ACTTGAGCTT	CTCTATTATA	AACAGGTTGT	1860
GGGAGTTGCG	CATGCACTTA	ACCTTTTGTT	TGGGGTGGAG	TTATCCACTG	GAGTGTTTTT	1920
GGCCGCCATG	GATGCGTTTT	TATTTCTGTG	TTTCGCCTCT	TTCCTTGTTAG	TTACTTACAA	1980
TTCTTTGCTG	TTCTTAATTT	TTTTATTATG	TAGTAAAATT	TTGATTCCTC	TGACTTGAGC	2040
TTCTCTATTA	TAAACAGGAA	AATGGTATGG	CAAATACAGT	ATCCATTTAC	TCTGCAGGCC	2100
TGGTATTACT	TCTCTATGTA	TCTGGCGTCT	TGCTGAGTCA	GTCTGAGATC	CCACTCTCTA	2160
TGAATGGAGT	GTTAACTCGG	TTAAATGGAG	AGAGCGCATT	CGCACTGATG	GGTCTTCTTG	2220

GCGCAAGCAT CGTCCCTCAC AATTTTATA TCCATTCTTA TTTTGCTGGG GTACCTTTTT	2280
TCTCTTTATA TGTATCTCTC TTCTCTGTTA AGAAGCAATA ATTATACTAA GCAGTGAACG	2340
CTCTATTACA GGAAAGTACA TCTTCGTCTG ATGTCGACAA GAGCAGCTTG TGTCAAGACC	2400
ATTTGTTTCGC CATCTTTGGT GTCTTCAGCG GACTGTCACT TGTAATTAT GTATTGATGA	2460
ATGCAGCAGC TAATGTGTTT CACAGTACTG GCCTTGTTGGT ACTGACTTTT CACGATGCCT	2520
TGTCACTAAT GGAGCAGGTT TGTCTGACG GTTTTATGTT CGTATTAGTC AATAATTCAT	2580
TTTTAGGGAA AATGTTGAGA AATCTCTCGT GATTATTAAT TATCTTGTTT TTGATTGTTG	2640
ATCACAGGTA TTTATGAGTC CGCTCATTCC AGTGGTCTTT TTGATGCTCT TGTCTTCTC	2700
TAGTCAAATT ACCGCACTAG CTTGGGCTTT CGGTGGAGAG GTCGTCCTGC ATGACTTCCT	2760
GAAGATAGAA ATACCCGCTT GGCTTCATCG TGCTACAATC AGAATTCTTG CAGTTGCTCC	2820
TGCGCTTTAT TGTGTATGGA CATCTGGTGC AGACGGAATA TACCAGTTAC TTATATTCAC	2880
CCAGGTCTTG GTGGCAATGA TGCTTCCTTG CTCGGTAATA CCGCTTTTCC GCATTGCTTC	2940
GTCGAGACAA ATCATGGGTG TCCATAAAAT CCCTCAGGTT GGCGAGTTCC TCGCACTTAC	3000
AACGTTTTTG GGATTTCTGG GGTGAATGT TGTMTTGTT GTTGAGATGG TATTTGGGAG	3060
CAGTGACTGG GCTGGTGGTT TGAGATGGAA TACCGGTATG GGCACCTCGA TTCAGTACAC	3120
CACTCTGCTT GTATCGTCAT GTGCATCCTT ATGCCTGATA CTCTGGCTGG CAGCCACGCC	3180
GCTGAAATCT GCGAGTAACA GAGCGGAAGC TCAAATATGG AACATGGATG CTCAAAATGC	3240
TTTATCTTAT CCATCTGTTC AAGAAGAGGA AATTGAAAGA ACAGAAACAA GGAGGAACGA	3300
AGACGAATCA ATAGTGCAGT TGGAAAGCAG GGTAAAGGAT CAGTTGGATA CTACGTCTGT	3360
TACTAGCTCG GTCTATGATT TGCCAGAGAA CATTCTAATG ACGGATCAAG AAATCCGTTC	3420
GAGCCCTCCA GAGGAAAGAG AGTTGGATGT AAAGTACTCT ACCTCTCAAG TTAGTAGTCT	3480
TAAGGAAGAC TCTGATGTAA AGGAACAGTC TGTATTGCAG TCAACAGTGG TTAATGAGGT	3540
CAGTGATAAG GATCTGATTG TTGAAACAAA GATGGCGAAA ATTGAACCAA TGAGTCCTGT	3600
GGAGAAGATT GTTAGCATGG AGAATAACAG CAAGTTTATT GAAAAGGATG TTGAAGGGGT	3660
TTCATGGGAA ACAGAAGAAG CTACCAAAGC TGCTCCTACA AGCAACTTTA CTGTCGGATC	3720
TGATGGTCCT CTTTCATTCC GCAGCTTAAG TGGGGAAGGG GGAAGTGGGA CTGGAAGCCT	3780
TTCACGGTTG CAAGGTTTGG GACGTGCTGC CCGGAGACAC TTATCTGCGA TCCTTGATGA	3840
ATTTTGGGGA CATTTATATG ATTTTCATGG GCAATTGGTT GCTGAAGCCA GGGCAAAGAA	3900
ACTAGATCAG CTGTTTGGCA CTGATCAAAA GTCAGCCTCT TCTATGAAAG CAGATTCTGT	3960
TGGAAAAGAC ATTAGCAGTG GATATTGCAT GTCACCAACT GCGAAGGGAA TGGATTACAA	4020
GATGACTTCA AGTTTATATG ATTCAGTGAA GCAGCAGAGG ACACCGGGAA GTATCGATTG	4080
GTTGTATGGA TTACAAAGAG GTTCGTCACC GTCACCGTTG GTCAACCGTA TGCAGATGTT	4140
GGGTGCATAT GGTAACACCA CTAATAATAA TAATGCTTAC GAATTGAGTG AGAGAAGATA	4200
CTCTAGCCTG CGTGCTCCAT CATCTTCAGA GGGTTGGGAA CACCAACAAC CAGCTACAGT	4260

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

CTTTCTCTCTC TCTATCTCTA TCTCTCGTAG CTTGATAAGA GTTCTCTCTCT TTTGAAGATC	60
CGTTTCTCTCTC TCTCTCACTG AGACTATTGT TGTTAGGTCA ACTTGCATC ATGGCGATT	120
CGAAGGTCTG AAGCTGATTT CGAATGGTTT GGAGATATCC GTAGTGGTTA AGCATATGGA	180
AGTCTATGTT CTGCTCTTGG TTGCTCTGTT AGGGCTTCCT CCATTTGGAC CAACTTAGCT	240
GAATGTTGTA TGATCTCTCT CTTGAAGCA GCAAATAAGA AGAAGGTCTG GTCCTTAACT	300
TAACATCTGG TTAGTAGAGG AAACCTCAGC TATTATTAGG TAAAGAAAGA CTGTACAGAG	360
TTGTATAACA AGTAAGCGTT AGAGTGGCTT TGTTTGCCTC GGTGATAGAA GAACCGACTG	420
ATTCGTTGTT GTGTGTTAGC TTTGGAGGGA ATCAGATTTC GCGAGGGAAG GTGTTTTAGA	480
TCAAATCTGT GAATTTTACT CAACTGAGGC TTTTAGTGAA CCACGACTGT AGAGTTGACC	540
TTGAATCCTA CTCTGAGTAA TTATATTATC AGATAGATTT AGGATGGAAG CTGAAATTGT	600
GAATGTGAGA CCTCAGCTAG GGTTTATCCA GAGAATGGTT CCTGCTCTAC TTCCTGTCCT	660
TTTGGTTTCT GTCGGATATA TTGATCCCGG GAAATGGGTT GCAAATATCG AAGGAGGTGC	720
TCGTTTCGGG TATGACTTGG TGGCAATTAC TCTGCTTTTC AATTTTGCCG CCATCTTATG	780
CCAATATGTT GCAGCTCGCA TAAGCGTTGT GACTGGTAAA CACTTGGCTC AGATCTGCAA	840
TGAAGAATAT GACAAGTGA CGTGCAATGTT CTTGGGCATT CAGGCGGAGT TCTCAGCAAT	900
TCTGCTCGAC CTTACCATGG TTGTGGGAGT TGCGCATGCA CTTAACCTTT TGTTTGGGGT	960
GGAGTTATCC ACTGGAGTGT TTTTGGCCGC CATGGATGCG TTTTATTTC CTGTTTTGCG	1020
CTCTTTCTTT GAAAATGGTA TGGCAAATAC AGTATCCATT TACTCTGCAG GCCTGGTATT	1080
ACTTCTCTAT GTATCTGGCG TCTTGCTGAG TCAGTCTGAG ATCCCACTCT CTATGAATGG	1140
AGTGTTAACT CGGTTAAATG GAGAGAGCGC ATTCGCACTG ATGGGTCTTC TTGGCGCAAG	1200
CATCGTCCCT CACAATTTTT ATATCCATTC TTATTTTGCT GGGGAAAGTA CATCTTCGTC	1260
TGATGTCGAC AAGAGCAGCT TGTGTCAAGA CCATTTGTTT GCCATCTTTG GTGTCTTCAG	1320
CGGACTGTCA CTTGTAAATT ATGTATTGAT GAATGCAGCA GCTAATGTGT TTCACAGTAC	1380
TGGCCTTGTT GTACTGACTT TTCACGATGC CTTGTCACTA ATGGAGCAGG TATTTATGAG	1440
TCCGCTCATT CCAGTGGTCT TTTTGATGCT CTTGTTCTTC TCTAGTCAA TTACCGCACT	1500
AGCTTGGGCT TTCGGTGGAG AGGTCGTCCT GCATGACTTC CTGAAGATAG AAATACCCGC	1560
TTGGCTTCAT CGTGCTACAA TCAGAATTCT TGCAGTTGCT CCTGCGCTTT ATTGTGTATG	1620
GACATCTGGT GCAGACGGAA TATACCAGTT ACTTATATTC ACCCAGGTCT TGGTGGCAAT	1680
GATGCTTCCT TGCTCGGTAA TACCGCTTTT CCGCATTGCT TCGTCGAGAC AAATCATGGG	1740

TGTCCATAAA	ATCCCTCAGG	TTGGCGAGTT	CCTCGCACTT	ACAACGTTTT	TGGGATTTCT	1800
GGGGTTGAAT	GTTGTTTTTG	TTGTTGAGAT	GGTATTTGGG	AGCAGTGAAT	GGGCTGGTGG	1860
TTTGAGATGG	AATACCGGTA	TGGGCACCTC	GATTCAGTAC	ACCACTCTGC	TTGTATCGTC	1920
ATGTGCATCC	TTATGCCTGA	TACTCTGGCT	GGCAGCCACG	CCGCTGAAAT	CTGCGAGTAA	1980
CAGAGCGGAA	GCTCAAATAT	GGAACATGGA	TGCTCAAAAT	GCTTTATCTT	ATCCATCTGT	2040
TCAAGAAGAG	GAAATTGAAA	GAACAGAAAC	AAGGAGGAAC	GAAGACGAAT	CAATAGTGCG	2100
GTTGGAAAGC	AGGGTAAAGG	ATCAGTTGGA	TACTACGTCT	GTTACTAGCT	CGGTCTATGA	2160
TTTGCCAGAG	AACATTCTAA	TGACGGATCA	AGAAATCCGT	TCGAGCCCTC	CAGAGGAAAG	2220
AGAGTTGGAT	GTAAAGTACT	CTACCTCTCA	AGTTAGTAGT	CTTAAGGAAG	ACTCTGATGT	2280
AAAGGAACAG	TCTGTATTGC	AGTCAACAGT	GGTTAATGAG	GTCAGTGATA	AGGATCTGAT	2340
TGTTGAAACA	AAGATGGCGA	AAATTGAACC	AATGAGTCCT	GTGGAGAAGA	TTGTTAGCAT	2400
GGAGAATAAC	AGCAAGTTTA	TTGAAAAGGA	TGTTGAAGGG	GTTTCATGGG	AAACAGAAGA	2460
AGCTACCAAA	GCTGCTCCTA	CAAGCAACTT	TACTGTCCGA	TCTGATGGTC	CTCCTTCATT	2520
CCGCAGCTTA	AGTGGGGAAG	GGGGAAGTGG	GACTGGAAGC	CTTTCACGGT	TGCAAGGTTT	2580
GGGACGTGCT	GGCCGGAGAC	ACTTATCTGC	GATCCTTGAT	GAATTTTGGG	GACATTTATA	2640
TGATTTTCAT	GGGCAATTGG	TTGCTGAAGC	CAGGGCAAAG	AAACTAGATC	AGCTGTTTGG	2700
CACTGATCAA	AAGTCAGCCT	CTTCTATGAA	AGCAGATTCG	TTTGGAAAAG	ACATTAGCAG	2760
TGGATATTGC	ATGTCACCAA	CTGCGAAGGG	AATGGATTCA	CAGATGACTT	CAAGTTTATA	2820
TGATTCAC TG	AAGCAGCAGA	GGACACCGGG	AAGTATCGAT	TCGTTGTATG	GATTACAAAG	2880
AGGTTTCGTC	CCGTCACCGT	TGGTCAACCG	TATGCAGATG	TTGGGTGCAT	ATGGTAACAC	2940
CACTAATAAT	AATAATGCTT	ACGAATTGAG	TGAGAGAAGA	TACTCTAGCC	TGCGTGCTCC	3000
ATCATCTTCA	GAGGGTTGGG	AACACCAACA	ACCAGCTACA	GTTACCGGAT	ACCAGATGAA	3060
GTCATATGTA	GACAATTGCG	CAAAAGAAAG	GCTTGAAGCC	TTACAATCCC	GTGGAGAGAT	3120
CCCACATCG	AGATCTATGG	CGCTTGGTAC	ATTGAGCTAT	ACACAGCAAC	TTGCTTTAGC	3180
CTTGAAACAG	AAGTCCCAGA	ATGGTCTAAC	CCCTGGACCA	GCTCCTGGGT	TTGAGAATTT	3240
TGCTGGGTCT	AGAAGCATAT	CGCGACAATC	TGAAAGATCT	TATTACGGTG	TTCCATCTTC	3300
TGGCAATACT	GATACTGTTG	GCGCAGCAGT	AGCCAATGAG	AAAAAATATA	GTAGCATGCC	3360
AGATATCTCA	GGATTGTCTA	TGTCCGCAAG	GAACATGCAT	TTACCAAACA	ACAAGAGTGG	3420
ATACTGGGAT	CCGTCAAGTG	GAGGAGGAGG	GTATGGTGCG	TCTTATGGTC	GGTTAAGCAA	3480
TGAATCATCG	TTATATTCTA	ATTTGGGGTC	ACGGGTGGGA	GTACCCTCGA	CTTATGATGA	3540
CATTTCTCAA	TCAAGAGGAG	GCTACAGAGA	TGCCTACAGT	TTGCCACAGA	GTGCAACAAC	3600
AGGGACCGGA	TCGCTTTGGT	CCAGACAGCC	CTTTGAGCAG	TTTGGTGTAG	CGGAGAGGAA	3660
TGGTGCTGTT	GGTGAGGAGC	TCAGGAATAG	ATCGAATCCG	ATCAATATAG	ACAACAACGC	3720
TTCTTCTAAT	GTTGATGCAG	AGGCTAAGCT	TCTTCAGTCG	TTCAGGCACT	GTATTCTAAA	3780

GCTTATTAAA CTTGAAGGAT CCGAGTGGTT GTTTGGACAA AGCGATGGAG TTGATGAAGA 3840
 ACTGATTGAC CGGGTAGCTG CACGAGAGAA GTTTATCTAT GAAGCTGAAG CTCGAGAAAT 3900
 AAACCAGGTG GGTACATGG GGGAGCCACT AATTTTCATCG GTTCCTAACT GTGGAGATGG 3960
 TTGCGTTTGG AGAGCTGATT TGATTGTGAG CTTTGGAGTT TGGTGCATTG ACCGTGTCCT 4020
 TGACTTGTCT CTCATGGAGA GTCGGCCTGA GCTTTGGGGA AAGTACACTT ACGTTCTCAA 4080
 CCGCTACAG GGAGTGATTG ATCCGGCGTT CTCAAAGCTG CGGACACCAA TGACACCGTG 4140
 CTTTTGCCTT CAGATTCCAG CGAGCCACCA GAGAGCGAGT CCGACTTCAG CTAACGGAAT 4200
 GTTACCTCCG GCTGCAAAAC CGGCTAAAGG CAAATGCACA ACCGCAGTCA CACTTCTTGA 4260
 TCTAATCAAA GACGTTGAAA TGGCAATCTC TTGTAGAAAA GGCCGAACCG GTACAGCTGC 4320
 AGGTGATGTG GCTTTCCCAA AGGGGAAAGA GAATTTGGCT TCGGTTTCGA AGCGGTATAA 4380
 ACGTCGGTTA TCGAATAAAC CAGTAAGGTA TGAATCAGGA TGGACCCGGT TCAAGAAAAA 4440
 ACGTGACTGC GTACGGATCA TTGGGTTGAA GAAGAAGAAC ATTGTGAGAA ATCTCATGAT 4500
 CAAAGTGACG TCGAGAGGGA AGCCGAAGAA TCAAACTCT CGCTTTTGAT TGCTCCTCTG 4560
 CTTGTTAAT TGTGTATTAA GAAAAGAAGA AAAAAATGG ATTTTGTGTT CTCAGAATT 4620
 TTTGCTCTT TTTTCTTAA TTGGTTGTA ATGTTATGTT TATATACATA TATCATCATC 4680
 ATAGGACCAT AGCTACAAAC CGAATCCGGT TTGTGTAATT CTATGCGGAA TCATAAGAA 4740
 ATCGTCG 4747

(2) INFORMATION FOR SEQ ID NO:3:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1321 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Met Glu Ala Glu Ile Val Asn Val Arg Pro Gln Leu Gly Phe Ile Gln
 1 5 10 15
 Arg Met Val Pro Ala Leu Leu Pro Val Leu Leu Val Ser Val Gly Tyr
 20 25 30
 Ile Asp Pro Gly Lys Trp Val Ala Asn Ile Glu Gly Gly Ala Arg Phe
 35 40 45
 Gly Tyr Asp Leu Val Ala Ile Thr Leu Leu Phe Asn Phe Ala Ala Ile
 50 55 60
 Leu Cys Gln Tyr Val Ala Ala Arg Ile Ser Val Val Thr Gly Lys His
 65 70 75 80

Leu	Ala	Gln	Ile	Cys 85	Asn	Glu	Glu	Tyr	Asp 90	Lys	Trp	Thr	Cys	Met 95	Phe
Leu	Gly	Ile	Gln 100	Ala	Glu	Phe	Ser	Ala 105	Ile	Leu	Leu	Asp	Leu 110	Thr	Met
Val	Val	Gly 115	Val	Ala	His	Ala	Leu 120	Asn	Leu	Leu	Phe	Gly 125	Val	Glu	Leu
Ser	Thr 130	Gly	Val	Phe	Leu	Ala 135	Ala	Met	Asp	Ala	Phe 140	Leu	Phe	Pro	Val
Phe 145	Ala	Ser	Phe	Leu	Glu 150	Asn	Gly	Met	Ala	Asn 155	Thr	Val	Ser	Ile	Tyr 160
Ser	Ala	Gly	Leu	Val 165	Leu	Leu	Leu	Tyr	Val 170	Ser	Gly	Val	Leu	Leu	Ser 175
Gln	Ser	Glu	Ile 180	Pro	Leu	Ser	Met	Asn 185	Gly	Val	Leu	Thr	Arg 190	Leu	Asn
Gly	Glu	Ser 195	Ala	Phe	Ala	Leu	Met 200	Gly	Leu	Leu	Gly	Ala 205	Ser	Ile	Val
Pro	His 210	Asn	Phe	Tyr	Ile	His 215	Ser	Tyr	Phe	Ala	Gly 220	Glu	Ser	Thr	Ser
Ser 225	Ser	Asp	Val	Asp	Lys 230	Ser	Ser	Leu	Cys	Gln 235	Asp	His	Leu	Phe	Ala 240
Ile	Phe	Gly	Val	Phe 245	Ser	Gly	Leu	Ser	Leu 250	Val	Asn	Tyr	Val	Leu	Met 255
Asn	Ala	Ala	Ala 260	Asn	Val	Phe	His	Ser 265	Thr	Gly	Leu	Val	Val	Leu	Thr 270
Phe	His	Asp 275	Ala	Leu	Ser	Leu	Met 280	Glu	Gln	Val	Phe	Met 285	Ser	Pro	Leu
Ile	Pro 290	Val	Val	Phe	Leu	Met 295	Leu	Leu	Phe	Phe	Ser 300	Ser	Gln	Ile	Thr
Ala 305	Leu	Ala	Trp	Ala	Phe 310	Gly	Gly	Glu	Val	Val 315	Leu	His	Asp	Phe	Leu 320
Lys	Ile	Glu	Ile	Pro 325	Ala	Trp	Leu	His	Arg 330	Ala	Thr	Ile	Arg	Ile	Leu 335
Ala	Val	Ala	Pro 340	Ala	Leu	Tyr	Cys	Val 345	Trp	Thr	Ser	Gly	Ala 350	Asp	Gly
Ile	Tyr	Gln 355	Leu	Leu	Ile	Phe	Thr 360	Gln	Val	Leu	Val	Ala 365	Met	Met	Leu
Pro	Cys 370	Ser	Val	Ile	Pro	Leu 375	Phe	Arg	Ile	Ala	Ser 380	Ser	Arg	Gln	Ile
Met 385	Gly	Val	His	Lys	Ile 390	Pro	Gln	Val	Gly	Glu 395	Phe	Leu	Ala	Leu	Thr 400
Thr	Phe	Leu	Gly	Phe 405	Leu	Gly	Leu	Asn	Val 410	Val	Phe	Val	Val	Glu	Met 415
Val	Phe	Gly	Ser 420	Ser	Asp	Trp	Ala	Gly 425	Gly	Leu	Arg	Trp	Asn 430	Thr	Gly
Met	Gly	Thr	Ser	Ile	Gln	Tyr	Thr	Thr	Leu	Leu	Val	Ser	Ser	Cys	Ala

435					440					445					
Ser	Leu	Cys	Leu	Ile	Leu	Trp	Leu	Ala	Ala	Thr	Pro	Leu	Lys	Ser	Ala
450					455					460					
Ser	Asn	Arg	Ala	Glu	Ala	Gln	Ile	Trp	Asn	Met	Asp	Ala	Gln	Asn	Ala
465				470					475					480	
Leu	Ser	Tyr	Pro	Ser	Val	Gln	Glu	Glu	Glu	Ile	Glu	Arg	Thr	Glu	Thr
				485					490					495	
Arg	Arg	Asn	Glu	Asp	Glu	Ser	Ile	Val	Arg	Leu	Glu	Ser	Arg	Val	Lys
			500					505					510		
Asp	Gln	Leu	Asp	Thr	Thr	Ser	Val	Thr	Ser	Ser	Val	Tyr	Asp	Leu	Pro
		515					520					525			
Glu	Asn	Ile	Leu	Met	Thr	Asp	Gln	Glu	Ile	Arg	Ser	Ser	Pro	Pro	Glu
		530				535					540				
Glu	Arg	Glu	Leu	Asp	Val	Lys	Tyr	Ser	Thr	Ser	Gln	Val	Ser	Ser	Leu
		545				550					555				560
Lys	Glu	Asp	Ser	Asp	Val	Lys	Glu	Gln	Ser	Val	Leu	Gln	Ser	Thr	Val
				565					570					575	
Val	Asn	Glu	Val	Ser	Asp	Lys	Asp	Leu	Ile	Val	Glu	Thr	Lys	Met	Ala
			580					585					590		
Lys	Ile	Glu	Pro	Met	Ser	Pro	Val	Glu	Lys	Ile	Val	Ser	Met	Glu	Asn
		595					600					605			
Asn	Ser	Lys	Phe	Ile	Glu	Lys	Asp	Val	Glu	Gly	Val	Ser	Trp	Glu	Thr
		610					615				620				
Glu	Glu	Ala	Thr	Lys	Ala	Ala	Pro	Thr	Ser	Asn	Phe	Thr	Val	Gly	Ser
		625				630					635				640
Asp	Gly	Pro	Pro	Ser	Phe	Arg	Ser	Leu	Ser	Gly	Glu	Gly	Gly	Ser	Gly
				645					650					655	
Thr	Gly	Ser	Leu	Ser	Arg	Leu	Gln	Gly	Leu	Gly	Arg	Ala	Ala	Arg	Arg
			660					665					670		
His	Leu	Ser	Ala	Ile	Leu	Asp	Glu	Phe	Trp	Gly	His	Leu	Tyr	Asp	Phe
		675					680					685			
His	Gly	Gln	Leu	Val	Ala	Glu	Ala	Arg	Ala	Lys	Lys	Leu	Asp	Gln	Leu
		690				695					700				
Phe	Gly	Thr	Asp	Gln	Lys	Ser	Ala	Ser	Ser	Met	Lys	Ala	Asp	Ser	Phe
		705				710					715				720
Gly	Lys	Asp	Ile	Ser	Ser	Gly	Tyr	Cys	Met	Ser	Pro	Thr	Ala	Lys	Gly
				725					730					735	
Met	Asp	Ser	Gln	Met	Thr	Ser	Ser	Leu	Tyr	Asp	Ser	Leu	Lys	Gln	Gln
			740					745					750		
Arg	Thr	Pro	Gly	Ser	Ile	Asp	Ser	Leu	Tyr	Gly	Leu	Gln	Arg	Gly	Ser
		755					760					765			
Ser	Pro	Ser	Pro	Leu	Val	Asn	Arg	Met	Gln	Met	Leu	Gly	Ala	Tyr	Gly
		770				775					780				
Asn	Thr	Thr	Asn	Asn	Asn	Asn	Ala	Tyr	Glu	Leu	Ser	Glu	Arg	Arg	Tyr
				785		790					795				800

Ser Ser Leu Arg Ala Pro Ser Ser Ser Glu Gly Trp Glu His Gln Gln
 805 810 815
 Pro Ala Thr Val His Gly Tyr Gln Met Lys Ser Tyr Val Asp Asn Leu
 820 825 830
 Ala Lys Glu Arg Leu Glu Ala Leu Gln Ser Arg Gly Glu Ile Pro Thr
 835 840 845
 Ser Arg Ser Met Ala Leu Gly Thr Leu Ser Tyr Thr Gln Gln Leu Ala
 850 855 860
 Leu Ala Leu Lys Gln Lys Ser Gln Asn Gly Leu Thr Pro Gly Pro Ala
 865 870 875 880
 Pro Gly Phe Glu Asn Phe Ala Gly Ser Arg Ser Ile Ser Arg Gln Ser
 885 890 895
 Glu Arg Ser Tyr Tyr Gly Val Pro Ser Ser Gly Asn Thr Asp Thr Val
 900 905 910
 Gly Ala Ala Val Ala Asn Glu Lys Lys Tyr Ser Ser Met Pro Asp Ile
 915 920 925
 Ser Gly Leu Ser Met Ser Ala Arg Asn Met His Leu Pro Asn Asn Lys
 930 935 940
 Ser Gly Tyr Trp Asp Pro Ser Ser Gly Gly Gly Gly Tyr Gly Ala Ser
 945 950 955 960
 Tyr Gly Arg Leu Ser Asn Glu Ser Ser Leu Tyr Ser Asn Leu Gly Ser
 965 970 975
 Arg Val Gly Val Pro Ser Thr Tyr Asp Asp Ile Ser Gln Ser Arg Gly
 980 985 990
 Gly Tyr Arg Asp Ala Tyr Ser Leu Pro Gln Ser Ala Thr Thr Gly Thr
 995 1000 1005
 Gly Ser Leu Trp Ser Arg Gln Pro Phe Glu Gln Phe Gly Val Ala Glu
 1010 1015 1020
 Arg Asn Gly Ala Val Gly Glu Glu Leu Arg Asn Arg Ser Asn Pro Ile
 1025 1030 1035 1040
 Asn Ile Asp Asn Asn Ala Ser Ser Asn Val Asp Ala Glu Ala Lys Leu
 1045 1050 1055
 Leu Gln Ser Phe Arg His Cys Ile Leu Lys Leu Ile Lys Leu Glu Gly
 1060 1065 1070
 Ser Glu Trp Leu Phe Gly Gln Ser Asp Gly Val Asp Glu Glu Leu Ile
 1075 1080 1085
 Asp Arg Val Ala Ala Arg Glu Lys Phe Ile Tyr Glu Ala Glu Ala Arg
 1090 1095 1100
 Glu Ile Asn Gln Val Gly His Met Gly Glu Pro Leu Ile Ser Ser Val
 1105 1110 1115 1120
 Pro Asn Cys Gly Asp Gly Cys Val Trp Arg Ala Asp Leu Ile Val Ser
 1125 1130 1135
 Phe Gly Val Trp Cys Ile His Arg Val Leu Asp Leu Ser Leu Met Glu
 1140 1145 1150
 Ser Arg Pro Glu Leu Trp Gly Lys Tyr Thr Tyr Val Leu Asn Arg Leu

55

1155	1160	1165
Gln Gly Val Ile Asp Pro Ala Phe Ser Lys Leu Arg Thr Pro Met Thr 1170	1175	1180
Pro Cys Phe Cys Leu Gln Ile Pro Ala Ser His Gln Arg Ala Ser Pro 1185	1190	1195 1200
Thr Ser Ala Asn Gly Met Leu Pro Pro Ala Ala Lys Pro Ala Lys Gly 1205	1210	1215
Lys Cys Thr Thr Ala Val Thr Leu Leu Asp Leu Ile Lys Asp Val Glu 1220	1225	1230
Met Ala Ile Ser Cys Arg Lys Gly Arg Thr Gly Thr Ala Ala Gly Asp 1235	1240	1245
Val Ala Phe Pro Lys Gly Lys Glu Asn Leu Ala Ser Val Ser Lys Arg 1250	1255	1260
Tyr Lys Arg Arg Leu Ser Asn Lys Pro Val Arg Tyr Glu Ser Gly Trp 1265	1270	1275 1280
Thr Arg Phe Lys Lys Lys Arg Asp Cys Val Arg Ile Ile Gly Leu Lys 1285	1290	1295
Lys Lys Asn Ile Val Arg Asn Leu Met Ile Lys Val Thr Ser Arg Gly 1300	1305	1310
Lys Pro Lys Asn Gln Asn Ser Arg Phe 1315	1320	

(2) INFORMATION FOR SEQ ID NO:4:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 2310 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

TCTTCTTCTT CTTCCTCTTC CTCATCTCGT ATCTCTAACT TTTGTCGAAG TTCTTTTGAT	60
GAAACTAGGG TTTATTATCT TCTCCTTCTT TTTCCCATCA CCATAGAAAA GGCAGAGACC	120
TTTTTCTTCA TCATTTTAT TCTCCTTCTT CTTCTGCTGT TCATTTCTCC AGGTTACAAT	180
GATGTTTAAT GAGATGGGAA TGTGTGGAAA CATGGATTTC TTCTCTTCTG GATCACTTGG	240
TGAAGTTGAT TTCTGTCCTG TTCCACAAGC TGAGCCTGAT TCCATTGTTG AAGATGACTA	300
TACTGATGAT GAGATTGATG TTGATGAATT GGAGAGGAGG ATGTGGAGAG ACAAATGCG	360
GCTTAAACGT CTCAAGGAGC AGGATAAGGG TAAAGAAGGT GTTGATGCTG CTAAACAGAG	420
GCAGTCTCAA GAGCAAGCTA GGAGGAAGAA AATGTCTAGA GCTCAAGATG GGATCTTGAA	480

SUBSTITUTE SHEET (RULE 26)

GTATATGTTG AAGATGATGG AAGTTTGTAAGCTCAAGGC TTTGTTTATG GGATTATTCC	540
GGAGAATGGG AAGCCTGTGA CTGGTGCTTC TGATAATTTA AGGGAGTGGT GGAAAGATAA	600
GGTTAGGTTT GATCGTAATG GTCCTGCGGC TATTACCAAG TATCAAGCGG AGAATAATAT	660
CCCGGGGATT CATGAAGGTA ATAACCCGAT TGGACCGACT CCTCATACCT TGCAAGAGCT	720
TCAAGACACG ACTCTTGGAT CGCTTTTGTC TGC GTTGATG CAACACTGTG ATCCTCCTCA	780
GAGACGTTTT CCTTTGGAGA AAGGAGTTCC TCCTCCGCGG TGGCCTAATG GGAAAGAGGA	840
TTGGTGGCCT CAACTTGGTT TGCCTAAAGA TCAAGGTCCT GCACCTTACA AGAAGCCTCA	900
TGATTTGAAG AAGGCGTGGA AAGTCGGCGT TTTGACTGCG GTTATCAAGC ATATGTTTTCC	960
TGATATTGCT AAGATCCGTA AGCTCGTGAG GCAATCTAAA TGTTTGCAGG ATAAGATGAC	1020
TGCTAAAGAG AGTGCTACCT GGCTTGCTAT TATTAACCAA GAAGAGTCCT TGGCTAGAGA	1080
GCTTTATCCC GAGTCATGTC CACCTCTTTC TCTGTCTGGT GGAAGTTGCT CGCTTCTGAT	1140
GAATGATTGC AGTCAATACG ATGTTGAAGG TTTGAGAGAG GAGTCTCACT ATGAAGTGGA	1200
AGAGCTCAAG CCAGAAAAAG TTATGAATTC TTCAAACCTT GGGATGGTTG CTAAAATGCA	1260
TGACTTTCCT GTCAAAGAAG AAGTCCCAGC AGGAAACTCG GAATTCATGA GAAAGAGAAA	1320
GCCAAACAGA GATCTGAACA CTATTATGGA CAGAACC GTT TTCACCTGCG AGAATCTTGG	1380
GTGTGCGCAC AGCGAAATCA GCCGGGGATT TCTGGATAGG AATTCGAGAG ACAACCATCA	1440
ACTGGCATGT CCACATCGAG ACAGTCGCTT ACCGTATGGA GCAGCACCAT CCAGGTTTTCA	1500
TGTCAATGAA GTTAAGCCTG TAGTTGGATT TCCTCAGCCA AGGCCAGTGA ACTCAGTAGC	1560
CCAACCAATT GACTTAACGG GTATAGTTCC TGAAGATGGA CAGAAGATGA TCTCAGAGCT	1620
CATGTCCATG TACGACAGAA ATGTCCAGAG CAACCAAACC TCTATGGTCA TGGAAAATCA	1680
AAGCGTGTCA CTGCTTCAAC CCACAGTCCA TAACCATCAA GAACATCTCC AGTTC CCAGG	1740
AAACATGGTG GAAGGAAGTT TCTTTGAAGA CTTGAACATC CCAAACAGAG CAAACAACAA	1800
CAACAGCAGC AACAATCAAA CGTTTTTTTCA AGGGAACAAC AACACAACA ATGTGTTTTAA	1860
GTTGACACT GCAGATCACA ACAACTTTGA AGCTGCACAT AACACAACA ATAACAGTAG	1920
CGGCAACAGG TTCCAGCTTG TGTGTGATTC CACACCGTTC GACATGGCGT CATTCGATTA	1980
CAGAGATGAT ATGTCGATGC CAGGAGTAGT AGGAACGATG GATGGAATGC AGCAGAAGCA	2040
GCAAGATGTA TCCATATGGT TCTAAAGTCT TGGTAGTAGA TTTCATCTTC TCTTATTTTT	2100
ATCTTTTGTG TTCTTACATT CACTCAACCA TGTAATATTT TTTCTGGGT CTCTCTGTCT	2160
CTATCGCTTG TTATGATGTG TCTGTAAGAG TCTCTAAAAA CTCTCTGTGA CTGTGTGTCT	2220
TTGTCTCGGC TTGGTGAATC TCTCTGTCAT CATCAGCTTT TAGTTACACA CCCGACTTGG	2280
GGATGAACGA AACTAAATG TAAGTTTTCA	2310

(2) INFORMATION FOR SEQ ID NO:5:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 3387 base pairs
- (B) TYPE: nucleic acid

(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

AGAGCAGTGA GTATTNCCAC NAGCCGCTTT GTTAATTACA TATTAATTGT GTAATAATAA	60
TAATAAATGA TGTCTTAAAT TTTATGTGTA AGAAATGAAA TTAAATGAT ATATATGTAT	120
ATTATATATC TANACATATA TATATATATA TAAATAGAGT ATATATACTA TGATCTATCT	180
TCCTGATCTA CAGAGAGACT CCACAAAGAA ACGCAAATAA ACAAAGTCG CTTTCTAGCC	240
ACGTGATCTT TCGTCGACTT TTCTTCTTCT TCTTCTTCTT CCTCTTCCTC ATCTCGTATC	300
TCTAACTTTT GTCGAAGTTC TTTTGATGAA ACTAGGGTTT ATTATCTTCT CCTTCTTTTT	360
CCCATCACCA TAGAAAAGGC AGAGACCTTT TTCTTCATCA TTTTATTCTT CCTTCTTCTT	420
CTGCTGTTCA TTTCTCCAGG TACTATACGC TTCTTCTTCT ATTGATTTTT TAGGGTTATT	480
ATTGATACTG AAGATGATGA TAGGTTTATT CATAGGGTTT TACTAGATCG ATGGTTTAC	540
TTTAGTTTAC TAGTGTTTAC ACGATCTAAT TTCATGAGTT TATNCTACTT TTAGTTTTTT	600
NTTGGGTGA AGTTTTGTTT ATTGTTTATA AATCGTTGAT CTATTTGAAA ATGTTTTCTC	660
TTTCTTATTC ATATATGATC CTTTCTATAT TTGGTTCCTA TGTTGAAGAT CTCATCCTTT	720
TTTTGGAAAT TGAATCTGTT GATAATTTTT ATTATCCGAT TGATTATTTA GTTTAGGAGT	780
GATTAAAATA CGATCTGATT ATGTGTTTAT TACTTAAAAC TTTGATTGAA TTCGAAAAGC	840
CCCTTTTTTA TAATTTAGGG TTTGATGATT TTTTITAGTA AGTTGTTTGA TTCAGAAGAA	900
ATATAATTGT ACTGATTAGT TTTGTTTGTG TATTTGATTT GTTACAGGTT ACAATGATGT	960
TTAATGAGAT GGAATGTGT GGAAACATGG ATTTCTTCTC TTCTGGATCA CTTGGTGAAG	1020
TTGATTTCTG TCCTGTTCCA CAAGCTGAGC CTGATTCCAT TGTTGAAGAT GACTATACTG	1080
ATGATGAGAT TGATGTTGAT GAATTGGAGA GGAGGATGTG GAGAGACAAA ATGCGGCTTA	1140
AACGTCTCAA GGAGCAGGAT AAGGGTAAAG AAGGTGTTGA TGCTGCTAAA CAGAGGCAGT	1200
CTCAAGAGCA AGCTAGGAGG AAGAAAATGT CTAGAGCTCA AGATGGGATC TTGAAGTATA	1260
TGTTGAAGAT GATGGAAGTT TGTAAGCTC AAGGCTTTGT TTATGGGATT ATTCCGAGA	1320
ATGGGAAGCC TGTGACTGGT GCTTCTGATA ATTTAAGGGA GTGGTGGAAA GATAAGGTTA	1380
GGTTTGATCG TAATGGTCCT GCGGCTATTA CCAAGTATCA AGCGGAGAAT AATATCCCGG	1440
GGATTCATGA AGGTAATAAC CCGATTGGAC CGACTCCTCA TACCTTGCAA GAGCTTCAAG	1500
ACACGACTCT TGGATCGCTT TTGTCTGCGT TGATGCAACA CTGTGATCCT CCTCAGAGAC	1560
GTTTTCTTTT GGAGAAAGGA GTTCCTCCTC CGTGGTGGCC TAATGGGAAA GAGGATTGGT	1620

GGCCTCAACT	TGGTTTGCCT	AAAGATCAAG	GTCCTGCACC	TTACAAGAA3	CCTCATGATT	1680
TGAAGAAGGC	GTGGAAAGTC	GGCGTTTTGA	CTGCGGTTAT	CAAGCATATG	TTTCCTGATA	1740
TTGCTAAGAT	CCGTAAGCTC	GTGAGGCAAT	CTAAATGTTT	GCAGGATAAG	ATGAC TGCTA	1800
AAGAGAGTGC	TACCTGGCTT	GCTATTATTA	ACCAAGAAGA	GTCCTTGGCT	AGAGAGCTTT	1860
ATCCCGAGTC	ATGTCCACCT	CTTCTCTGT	CTGGTGGAAG	TTGCTCGCTT	CTGATGAATG	1920
ATTGCAGTCA	ATACGATGTT	GAAGGTTTCG	AGAAGGAGTC	TCACTATGAA	GTGGAAGAGC	1980
TCAAGCCAGA	AAAAGTTATG	AATTCTTCAA	ACTTTGGGAT	GGTTGCTAAA	ATGCATGACT	2040
TTCTGTCAA	AGAAGAAGTC	CCAGCAGGAA	ACTCGGAATT	CATGAGAAAAG	AGAAAAGCCAA	2100
ACAGAGATCT	GAACACTATT	ATGGACAGAA	CCGTTTTTCAC	CTGCGAGAAT	CTTGGGTGTG	2160
CGCACAGCGA	AATCAGCCGG	GGATTTCTGG	ATAGGAATTC	GAGAGACAAC	CATCAACTGG	2220
CATGTCCACA	TCGAGACAGT	CGCTTACCGT	ATGGAGCAGC	ACCATCCAGG	TTTCATGTCA	2280
ATGAAGTTAA	GCCTGTAGTT	GGATTTCTCT	AGCCAAGGCC	AGTGAAGTCA	GTAGCCCAAC	2340
CAATTGACTT	AACGGGTATA	GTTCTGAAG	ATGGACAGAA	GATGATCTCA	GAGCTCATGT	2400
CCATGTACGA	CAGAAATGTC	CAGAGCAACC	AAACCTCTAT	GGTCATGGAA	AATCAAAGCG	2460
TGTCACTGCT	TCAACCCACA	GTCCATAACC	ATCAAGAACA	TCTCCAGTTC	CCAGGAAACA	2520
TGGTGGAAGG	AAGTTTCTTT	GAAGACTTGA	ACATCCCAA	CAGAGCAAAC	AACAACAACA	2580
GCAGCAACAA	TCAAACGTTT	TTTCAAGGGA	ACAACAACAA	CAACAATGTG	TTTAAGTTCTG	2640
AACTGCAGA	TCACAACAAC	TTTGAAGCTG	CACATAACAA	CAACAATAAC	AGTAGCGGCA	2700
ACAGGTTCCA	GCTTGTGTTT	GATTCCACAC	CGTTCGACAT	GGCGTCATTC	GATTACAGAG	2760
ATGATATGTC	GATGCCAGGA	GATAGTAGGAA	CGATGGATGG	AATGCAGCAG	AAGCAGCAAG	2820
ATGTATCCAT	ATGGTTCTAA	AGTCTTGGTA	GATAGATTTC	TCTTCTCTTA	TTTTTATCTT	2880
TTGTGTTCTT	ACATTCATCT	AACCATGTAA	TATTTTTTCC	TGGGTCTCTC	TGTCTCTATC	2940
GCTTGTTATG	ATGTGTCTGT	AAGAGTCTCT	AAAACTCTC	TGTTACTGTG	TGTCTTTGTC	3000
TCGGCTTGGT	GAATCTCTCT	GTCAATCATCA	GCTTTTAGTT	ACACACCCGA	CTTGGGGATG	3060
AACGAACACT	AAATGTAAGT	TTTCATAATA	TAAATATATT	TGNAAGCTCT	CTTCTTCTGT	3120
GTGTTTTGGT	TGAGTTTGAC	TTTACAATT	GAAAAGTTTG	GTGTAATTCA	CGCTAACTAC	3180
CTCAAAGTTA	GGGAATGGTG	GGATAATTAT	TTATTACAAT	TGTATTTGAT	GGATAACGTG	3240
CTTATCGCTA	GTGGCTCGCG	GGTAGCATTT	AAGCATGGGT	CAATGCTTGT	GTCTACGAGC	3300
TCGAGTGATC	GAGCACACAC	AATCCAATCC	GAACACAAAA	CAAGAAGAAA	AACAAAATAA	3360
GATCTTAGAT	GTAAGGNATT	CTTAAAT				3387

(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 628 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

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- (ii) MOLECULE TYPE: peptide
 (iii) HYPOTHETICAL: NO
 (iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Met	Met	Phe	Asn	Glu	Met	Gly	Met	Cys	Gly	Asn	Met	Asp	Phe	Phe	Ser	1	5	10	15
Ser	Gly	Ser	Leu	Gly	Glu	Val	Asp	Phe	Cys	Pro	Val	Pro	Gln	Ala	Glu	20	25	30	
Pro	Asp	Ser	Ile	Val	Glu	Asp	Asp	Tyr	Thr	Asp	Asp	Glu	Ile	Asp	Val	35	40	45	
Asp	Glu	Leu	Glu	Arg	Arg	Met	Trp	Arg	Asp	Lys	Met	Arg	Leu	Lys	Arg	50	55	60	
Leu	Lys	Glu	Gln	Asp	Lys	Gly	Lys	Glu	Gly	Val	Asp	Ala	Ala	Lys	Gln	65	70	75	80
Arg	Gln	Ser	Gln	Glu	Gln	Ala	Arg	Arg	Lys	Lys	Met	Ser	Arg	Ala	Gln	85	90	95	
Asp	Gly	Ile	Leu	Lys	Tyr	Met	Leu	Lys	Met	Met	Glu	Val	Cys	Lys	Ala	100	105	110	
Gln	Gly	Phe	Val	Tyr	Gly	Ile	Ile	Pro	Glu	Asn	Gly	Lys	Pro	Val	Thr	115	120	125	
Gly	Ala	Ser	Asp	Asn	Leu	Arg	Glu	Trp	Trp	Lys	Asp	Lys	Val	Arg	Phe	130	135	140	
Asp	Arg	Asn	Gly	Pro	Ala	Ala	Ile	Thr	Lys	Tyr	Gln	Ala	Glu	Asn	Asn	145	150	155	160
Ile	Pro	Gly	Ile	His	Glu	Gly	Asn	Asn	Pro	Ile	Gly	Pro	Thr	Pro	His	165	170	175	
Thr	Leu	Gln	Glu	Leu	Gln	Asp	Thr	Thr	Leu	Gly	Ser	Leu	Leu	Ser	Ala	180	185	190	
Leu	Met	Gln	His	Cys	Asp	Pro	Pro	Gln	Arg	Arg	Phe	Pro	Leu	Glu	Lys	195	200	205	
Gly	Val	Pro	Pro	Pro	Trp	Trp	Pro	Asn	Gly	Lys	Glu	Asp	Trp	Trp	Pro	210	215	220	
Gln	Leu	Gly	Leu	Pro	Lys	Asp	Gln	Gly	Pro	Ala	Pro	Tyr	Lys	Lys	Pro	225	230	235	240
His	Asp	Leu	Lys	Lys	Ala	Trp	Lys	Val	Gly	Val	Leu	Thr	Ala	Val	Ile	245	250	255	
Lys	His	Met	Phe	Pro	Asp	Ile	Ala	Lys	Ile	Arg	Lys	Leu	Val	Arg	Gln	260	265	270	
Ser	Lys	Cys	Leu	Gln	Asp	Lys	Met	Thr	Ala	Lys	Glu	Ser	Ala	Thr	Trp	275	280	285	
Leu	Ala	Ile	Ile	Asn	Gln	Glu	Glu	Ser	Leu	Ala	Arg	Glu	Leu	Tyr	Pro	290	295	300	

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Glu Ser Cys Pro Pro Leu Ser Leu Ser Gly Gly Ser Cys Ser Leu Leu
 305 310 315 320
 Met Asn Asp Cys Ser Gln Tyr Asp Val Glu Gly Phe Glu Lys Glu Ser
 325 330 335
 His Tyr Glu Val Glu Glu Leu Lys Pro Glu Lys Val Met Asn Ser Ser
 340 345 350
 Asn Phe Gly Met Val Ala Lys Met His Asp Phe Pro Val Lys Glu Glu
 355 360 365
 Val Pro Ala Gly Asn Ser Glu Phe Met Arg Lys Arg Lys Pro Asn Arg
 370 375 380
 Asp Leu Asn Thr Ile Met Asp Arg Thr Val Phe Thr Cys Glu Asn Leu
 385 390 395 400
 Gly Cys Ala His Ser Glu Ile Ser Arg Gly Phe Leu Asp Arg Asn Ser
 405 410 415
 Arg Asp Asn His Gln Leu Ala Cys Pro His Arg Asp Ser Arg Leu Pro
 420 425 430
 Tyr Gly Ala Ala Pro Ser Arg Phe His Val Asn Glu Val Lys Pro Val
 435 440 445
 Val Gly Phe Pro Gln Pro Arg Pro Val Asn Ser Val Ala Gln Pro Ile
 450 455 460
 Asp Leu Thr Gly Ile Val Pro Glu Asp Gly Gln Lys Met Ile Ser Glu
 465 470 475 480
 Leu Met Ser Met Tyr Asp Arg Asn Val Gln Ser Asn Gln Thr Ser Met
 485 490 495
 Val Met Glu Asn Gln Ser Val Ser Leu Leu Gln Pro Thr Val His Asn
 500 505 510
 His Gln Glu His Leu Gln Phe Pro Gly Asn Met Val Glu Gly Ser Phe
 515 520 525
 Phe Glu Asp Leu Asn Ile Pro Asn Arg Ala Asn Asn Asn Asn Ser Ser
 530 535 540
 Asn Asn Gln Thr Phe Phe Gln Gly Asn Asn Asn Asn Asn Val Phe
 545 550 555 560
 Lys Phe Asp Thr Ala Asp His Asn Asn Phe Glu Ala Ala His Asn Asn
 565 570 575
 Asn Asn Asn Ser Ser Gly Asn Arg Phe Gln Leu Val Phe Asp Ser Thr
 580 585 590
 Pro Phe Asp Met Ala Ser Phe Asp Tyr Arg Asp Asp Met Ser Met Pro
 595 600 605
 Gly Val Val Gly Thr Met Asp Gly Met Gln Gln Lys Gln Gln Asp Val
 610 615 620
 Ser Ile Trp Phe
 625

(2) INFORMATION FOR SEQ ID NO:7:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2234 base pairs

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(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

GGCCGCTTCA AACTCTACAA ACCCAGAAAC CACCACACAG TAATTAATGT CTCCTTCTTT	60
CTTCCCATGT GATCTTTAAC AGACTTTTCT TCTTATTCTC CATCTCTGAA GTTGTGGGGA	120
TTCATCAAGA CTTCTTATC TGTTTCTTTT ATAAAACAAG AGAGAGATAC CACTTTTGGT	180
GTTCTTTATT TGCAACTCTT TCAGGTTAAA GAAATCGATA GGCTCTGTTC TTGATTGTGG	240
TGGAAGAGAC ATGATGATGT TTAACGAGAT GGAATGTAT GGAAACATGG ATTTCTTCTC	300
TTCCTCCACA TCTCTCGATG TGTGTCCATT ACCACAAGCT GAACAAGAAC CTGTAGTTGA	360
AGATGTCGAC TACACCGATG ATGAGATGGA TGAGCTTGAG CAGAGGATGT GGAGAGACAA	420
AATGCGTTTG AAACGTCTCA AGGAGCAACA GAGTAAGTGT AAAGGAGGCG TCGATGGTTC	480
GAAACAGAGG CAGTCGCAAG AGCAAGCTAG GAGGAAGAAA ATGTCTAGAG CCCAAGATGG	540
GATCTTGAAG TATATGTTGA AGATGATGGA AGTTTGTAAG GCTCAAGGCT TTGTTTATGG	600
TATTATTCCT GAGAAGGGTA AGCCTGTGAC TGGTGCTTCG GATAATTGTA GGGAAATGGT	660
GAAAGATAAG GTTAGGTTTG ATCGTAATGG TCCAGCTGCT ATTGCTAAGT ATCAGTCAGA	720
GAATAATATT TCTGGAGGGA GTAATGATTG TAACAGCTTG GTTGGTCCAA CACCGCATAC	780
GCTTCAGGAG CTTCAGGACA CGACTCTTGG TTCGCTTTTA TCGGCTTTGA TGCAACATTG	840
TGATCCACCG CAGAGACGGT TTCCTTTGGA GAAAGGAGTT TCTCCACCTT GGTGGCCTAA	900
TGGGAATGAA GAGTGGTGGC CTCAGCTTGG TTTACCAAAT GAGCAAGGTC CTCCTCCTTA	960
TAAGAAGCCT CATGATTGTA AGAAAGCTTG GAAAGTCGGT GTTTTAACTG CGGTGATCAA	1020
GCATATGTCG CCGGATATTG CGAAGATCCG TAAGCTTGTG AGGCAATCAA AATGCTTGCA	1080
GGATAAGATG ACGGCGAAAG AGAGTGCTAC TTGGCTTGCC ATTATTAACC AAGAAGAGGT	1140
TGTGGCTCGG GAGCTTTATC CCGAGTCATG CCCTCCTCTT TCTTCTTCTT CATCATTAGG	1200
AAGCGGGTCG CTTCTCATTG ATGATTGTAG CGAGTATGAC GTTGAAGGTT TCGAGAAGGA	1260
ACAACATGGT TTCGATGTGG AAGAGCGGAA ACCAGAGATA GTGATGATGC ATCCTCTAGC	1320
AAGCTTTGGG GTTGCTAAAA TGCAACATTT TCCCATAAAG GAGGAGGTCG CCACCACGGT	1380
AACTTAGAG TTCACGAGAA AGAGGAAGCA GAACAATGAT ATGAATGTTA TGGAATGGA	1440
CAGATCAGCA GGTTACACTT GTGAGAATGG TCAGTGTCTT CACAGCAAAA TGAATCTTGG	1500
ATTTCAAGAC AGGAGTTCAA GGGACAACCA CCAGATGGTT TGTCCATATA GAGACAATCG	1560
TTTAGCGTAT GGAGCATCCA AGTTTCATAT GGGTGAATG AACTAGTAG TTCCTCAGCA	1620

ACCAGTCCAA CCGATCGACC TATCGGGCGT TGGAGTTCCG GAAAACGGGC AGAAGATGAT 1680
 CACCGAGCTT ATGGCCATGT ACGACAGAAA TGTCCAAAGC AACCAAACGC CTCCTACTTT 1740
 GATGGAAAAC CAAAGCATGG TCATTGATGC AAAAGCAGCT CAGAATCAGC AGCTGAATTT 1800
 CAACAGTGGC AATCAAATGT TTATGCAACA AGGGACGAAC AACGGGGTTA ACAATCGGTT 1860
 CCAGATGGTG TTTGATTCTGA CACCATTCTGA TATGGCAGCA TTCGATTACA GAGATGATTG 1920
 GCAAACCGGA GCAATGGAAG GAATGGGGAA GCAGCAGCAG CAGCAGCAGC AGCAGCAAAG 1980
 ATGTATCAAT ATGGTTCTGA ATATTACACA ATCTCTGTAA TATTCATTCT TTCATAATAA 2040
 CTCTGTTACC TACTTACCTG ACTTGGGTAT GTATTCTATT GCACCAAACA CTCATCTATA 2100
 TTGTTGATGA TGATGAAGCC ATCTATTTTT TTTTGTGTC TGAAAGTCAT TTAACGCT 2160
 TCATTGTTTT AATAATGTCA CTATCCATTG AACATCATTC TCATGCTACA AGTTTGATTC 2220
 TTTGAGGCGG CCGC 2234

(2) INFORMATION FOR SEQ ID NO:8:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 584 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

Met Met Met Phe Asn Glu Met Gly Met Tyr Gly Asn Met Asp Phe Phe
 1 5 10 15
 Ser Ser Ser Thr Ser Leu Asp Val Cys Pro Leu Pro Gln Ala Glu Gln
 20 25 30
 Glu Pro Val Val Glu Asp Val Asp Tyr Thr Asp Asp Glu Met Asp Val
 35 40 45
 Asp Glu Leu Glu Lys Arg Met Trp Arg Asp Lys Met Arg Leu Lys Arg
 50 55 60
 Leu Lys Glu Gln Gln Ser Lys Cys Lys Glu Gly Val Asp Gly Ser Lys
 65 70 75 80
 Gln Arg Gln Ser Gln Glu Gln Ala Arg Arg Lys Lys Met Ser Arg Ala
 85 90 95
 Gln Asp Gly Ile Leu Lys Tyr Met Leu Lys Met Met Glu Val Cys Lys
 100 105 110
 Ala Gln Gly Phe Val Tyr Gly Ile Ile Pro Glu Lys Gly Lys Pro Val
 115 120 125
 Thr Gly Ala Ser Asp Asn Leu Arg Glu Trp Trp Lys Asp Lys Val Arg
 130 135 140

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Phe Asp Arg Asn Gly Pro Ala Ala Ile Ala Lys Tyr Gln Ser Glu Asn
 145 150 155 160
 Asn Ile Ser Gly Gly Ser Asn Asp Cys Asn Ser Leu Val Gly Pro Thr
 165 170 175
 Pro His Thr Leu Gln Glu Leu Gln Asp Thr Thr Leu Gly Ser Leu Leu
 180 185 190
 Ser Ala Leu Met Gln His Cys Asp Pro Pro Gln Arg Arg Phe Pro Leu
 195 200 205
 Glu Lys Gly Val Ser Pro Pro Trp Trp Pro Asn Gly Asn Glu Glu Trp
 210 215 220
 Trp Pro Gln Leu Gly Leu Pro Asn Glu Gln Gly Pro Pro Pro Tyr Lys
 225 230 235 240
 Lys Pro His Asp Leu Lys Lys Ala Trp Lys Val Gly Val Leu Thr Ala
 245 250 255
 Val Ile Lys His Met Ser Pro Asp Ile Ala Lys Ile Arg Lys Leu Val
 260 265 270
 Arg Gln Ser Lys Cys Leu Gln Asp Lys Met Thr Ala Lys Glu Ser Ala
 275 280 285
 Thr Trp Leu Ala Ile Ile Asn Gln Glu Glu Val Val Ala Arg Glu Leu
 290 295 300
 Tyr Pro Glu Ser Cys Pro Pro Leu Ser Ser Ser Ser Ser Leu Gly Ser
 305 310 315 320
 Gly Ser Leu Leu Ile Asn Asp Cys Ser Glu Tyr Asp Val Glu Gly Phe
 325 330 335
 Glu Lys Glu Gln His Gly Phe Asp Val Glu Glu Arg Lys Pro Glu Ile
 340 345 350
 Val Met Met His Pro Leu Ala Ser Phe Gly Val Ala Lys Met Gln His
 355 360 365
 Phe Pro Ile Lys Glu Glu Val Ala Thr Thr Val Asn Leu Glu Phe Thr
 370 375 380
 Arg Lys Arg Lys Gln Asn Asn Asp Met Asn Val Met Val Met Asp Arg
 385 390 395 400
 Ser Ala Gly Tyr Thr Cys Glu Asn Gly Gln Cys Pro His Ser Lys Met
 405 410 415
 Asn Leu Gly Phe Gln Asp Arg Ser Ser Arg Asp Asn His Gln Met Val
 420 425 430
 Cys Pro Tyr Arg Asp Asn Arg Leu Ala Tyr Gly Ala Ser Lys Phe His
 435 440 445
 Met Gly Gly Met Lys Leu Val Val Pro Gln Gln Pro Val Gln Pro Ile
 450 455 460
 Asp Leu Ser Gly Val Gly Val Pro Glu Asn Gly Gln Lys Met Ile Thr
 465 470 475 480
 Glu Leu Met Ala Met Tyr Asp Arg Asn Val Gln Ser Asn Gln Thr Pro
 485 490 495
 Pro Thr Leu Met Glu Asn Gln Ser Met Val Ile Asp Ala Lys Ala Ala

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500	505	510
Gln Asn Gln Gln Leu Asn Phe	Asn Ser Gly Asn Gln Met Phe Met Gln	
515	520	525
Gln Gly Thr Asn Asn Gly Val	Asn Asn Arg Phe Gln Met Val Phe Asp	
530	535	540
Ser Thr Pro Phe Asp Met Ala Ala Phe Asp Tyr Arg Asp Asp Trp Gln		
545	550	555
Thr Gly Ala Met Glu Gly Met Gly Lys Gln Gln Gln Gln Gln Gln		
565	570	575
Gln Gln Asp Val Ser Ile Trp Phe		
580		

(2) INFORMATION FOR SEQ ID NO:9:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 1722 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:

CAGATTCTAT GGATATGTAT AACACAATA TAGGGATGTT CCGGAGTTTA GTTTGTAGCT	60
CGGCGCCTCC ATTTACAGAG GGACATATGT GTTCTGATTC GCATACGGCT TTGTGCGATG	120
ATCTGAGTAG TGATGAGGAA ATGGAAATAG AGGAGCTTGA GAAGAAGATC TGGAGAGACA	180
AGCAGCGTTT AAAGCGGCTC AAGGAAATGG CGAAGAACGG TCTAGGAACA AGATTGTTGT	240
TGAAGCAGCA ACATGATGAT TTTCCAGAGC ACTCTAGTAA GAGAACCATG TACAAGGCAC	300
AAGATGGGAT CTTGAAGTAC ATGTCGAAGA CAATGGAGCG ATATAAAGCT CAAGGTTTTG	360
TTTATGGGAT TGTGTTAGAG AATGGGAAAA CGGTAGCGGG ATCTTCTGAT AATCTCCGTG	420
AATGGTGGAA AGACAAAGTG AGGTTTGATA GGAACGGCCC AGCTGCTATA ATCAAGCACC	480
AAAGGGATAT CAATCTTTCT GATGGAAGTG ATTCAGGGTC TGAGGTTGGG GATTCTACCG	540
CACAGAAGTT GCTTGAGCTT CAAGATACTA CTCTTGAGC TCTGTTATCG GCTCTGTTTC	600
CTCACTGCAA CCCTCCTCAG AGGCGGTTTC CGTTGGAGAA AGGCGTGACA CCGCCATGGT	660
GGCCAACGGG GAAAGAAGAT TGGTGGGATC AACTGTCTTT ACCCGTTGAT TTTGAGGTG	720
TTCCGCCACC TTACAAGAAG CCTCATGATC TCAAGAAGCT GTGGAAAATT GGTGTTTTGA	780
TTGGTGTAAT CAGACATATG GCTTCTGACA TTAGCAACAT ACCCAATCTC GTGAGACGGT	840
CTAGAAGTTT GCAGGAGAAA ATGACGTCAA GAGAAGGCGC TTTATGGCTC GCTGCTCTTT	900
ACCGAGAAAA GGCTATTGTT GATCAAATAG CCATGTCTAG AGAAAACAAC AACACTTCTA	960

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ACTTCTTGT TCCTGCAACC GGTGGAGACC CAGATGTTTT GTTTCCTGAA TCTACAGACT 1020
ATGATGTTGA ACTGATTGGT GGCACATCATC GGACCAATCA GCAGTATCCT GAATTTGAAA 1080
ACAACTACAA CTGTGTTTAC AAGAGAAAGT TTGAAGAAGA TTTTGGGATG CCAATGCATC 1140
CAACACTCCT AACATGTGAG AACAGTCTCT GTCCTTATAG CCAACCACAT ATGGGATTTC 1200
TTGACAGGAA CTTAAGAGAG AATCACCAAA TGACTTGTCC TTATAAAGTC ACTTCCTTCT 1260
ACCAACCAAC TAAACCCTAT GGTATGACGG GTTTAATGGT TCCTTGTCCG GATTATAACG 1320
GGATGCAGCA GCAGGTTTCA AGCTTTTCAAG ACCAGTTTAA TCATCCCAAC GATCTCTACA 1380
GACCAAAAGC TCCACAAAGA GGCAACGATG ACTTGTTTGA GGATTTGAAT CCTTCTCCTT 1440
CGACGCTGAA TCAGAATCTT GGTTTAGTCT TACCTACTGA CTTCAATGGA GGTGAGGAAA 1500
CAGTAGGAAC AGAGAACAAT CTGCATAATC AAGGGCAAGA GTTGCCCAACA TCTTGGATTC 1560
AGTAAAGAAA GCTTCAGAGT TTTCTTTTGA TGTTTTCTAG TCTTTATAGC TTTGTCTCTT 1620
GCTTATTCTC TCATTAAACA CAGTTTTTGA TCTCTCCATT TCATAGCCCA TGTAGCAATG 1680
GAGAAGATTA GGTTCATATA TAAGTTAATA ACCAAATTCA AA 1722

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(2) INFORMATION FOR SEQ ID NO:10:

- (i) SEQUENCE CHARACTERISTICS:
- (A) LENGTH: 520 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

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Asp Ser Met Asp Met Tyr Asn Asn Asn Ile Gly Met Phe Arg Ser Leu
1      5      10
Val Cys Ser Ser Ala Pro Pro Phe Thr Glu Gly His Met Cys Ser Asp
20     25     30
Ser His Thr Ala Leu Cys Asp Asp Leu Ser Ser Asp Glu Glu Met Glu
35     40     45
Ile Glu Glu Leu Glu Lys Lys Ile Trp Arg Asp Lys Gln Arg Leu Lys
50     55     60
Arg Leu Lys Glu Met Ala Lys Asn Gly Leu Gly Thr Arg Leu Leu Leu
65     70     75     80
Lys Gln Gln His Asp Asp Phe Pro Glu His Ser Ser Lys Arg Thr Met
85     90     95
Tyr Lys Ala Gln Asp Gly Ile Leu Lys Tyr Met Ser Lys Thr Met Glu
100    105    110
Arg Tyr Lys Ala Gln Gly Phe Val Tyr Gly Ile Val Leu Glu Asn Gly
115    120    125

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Lys Thr Val Ala Gly Ser Ser Asp Asn Leu Arg Glu Trp Trp Lys Asp
 130 135 140
 Lys Val Arg Phe Asp Arg Asn Gly Pro Ala Ala Ile Ile Lys His Gln
 145 150 155 160
 Arg Asp Ile Asn Leu Ser Asp Gly Ser Asp Ser Gly Ser Glu Val Gly
 165 170 175
 Asp Ser Thr Ala Gln Lys Leu Leu Glu Leu Gln Asp Thr Thr Leu Gly
 180 185 190
 Ala Leu Leu Ser Ala Leu Phe Pro His Cys Asn Pro Pro Gln Arg Arg
 195 200 205
 Phe Pro Leu Glu Lys Gly Val Thr Pro Pro Trp Trp Pro Thr Gly Lys
 210 215 220
 Glu Asp Trp Trp Asp Gln Leu Ser Leu Pro Val Asp Phe Arg Gly Val
 225 230 235 240
 Pro Pro Pro Tyr Lys Lys Pro His Asp Leu Lys Lys Leu Trp Lys Ile
 245 250 255
 Gly Val Leu Ile Gly Val Ile Arg His Met Ala Ser Asp Ile Ser Asn
 260 265 270
 Ile Pro Asn Leu Val Arg Arg Ser Arg Ser Leu Gln Glu Lys Met Thr
 275 280 285
 Ser Arg Glu Gly Ala Leu Trp Leu Ala Ala Leu Tyr Arg Glu Lys Ala
 290 295 300
 Ile Val Asp Gln Ile Ala Met Ser Arg Glu Asn Asn Asn Thr Ser Asn
 305 310 315 320
 Phe Leu Val Pro Ala Thr Gly Gly Asp Pro Asp Val Leu Phe Pro Glu
 325 330 335
 Ser Thr Asp Tyr Asp Val Glu Leu Ile Gly Gly Thr His Arg Thr Asn
 340 345 350
 Gln Gln Tyr Pro Glu Phe Glu Asn Asn Tyr Asn Cys Val Tyr Lys Arg
 355 360 365
 Lys Phe Glu Glu Asp Phe Gly Met Pro Met His Pro Thr Leu Leu Thr
 370 375 380
 Cys Glu Asn Ser Leu Cys Pro Tyr Ser Gln Pro His Met Gly Phe Leu
 385 390 395 400
 Asp Arg Asn Leu Arg Glu Asn His Gln Met Thr Cys Pro Tyr Lys Val
 405 410 415
 Thr Ser Phe Tyr Gln Pro Thr Lys Pro Tyr Gly Met Thr Gly Leu Met
 420 425 430
 Val Pro Cys Pro Asp Tyr Asn Gly Met Gln Gln Gln Val Gln Ser Phe
 435 440 445
 Gln Asp Gln Phe Asn His Pro Asn Asp Leu Tyr Arg Pro Lys Ala Pro
 450 455 460
 Gln Arg Gly Asn Asp Asp Leu Val Glu Asp Leu Asn Pro Ser Pro Ser
 465 470 475 480
 Thr Leu Asn Gln Asn Leu Gly Leu Val Leu Pro Thr Asp Phe Asn Gly

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														67																																	
														485																490																495	
Gly	Glu	Glu	Thr	Val	Gly	Thr	Glu	Asn	Asn	Leu	His	Asn	Gln	Gly	Gln																																
								500		505																510																					
Glu	Leu	Pro	Thr	Ser	Trp	Ile	Gln																																								
							515		520																																						

(2) INFORMATION FOR SEQ ID NO:11:

- (i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 2065 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

TTCCCTCTGAG	AACGACAGGA	GAAAGAATAA	AAACCCATAA	TTTCTTTAAT	TTCGGCGCTT	60
CAGATTATCG	TTGTTAAAGG	TTTTTGATTG	ATTTTGTTTA	AATGGGCGAT	CTTGCTATGT	120
CCGTAGCAGA	CATCAGGATG	GAGAATGAGC	CTGATGATTT	AGCTAGTGAT	AATGTTGCTG	180
AGATTGATGT	GAGTGATGAA	GAGATTGATG	CTGACGACCT	TGAGAGACGG	ATGTGGAAAG	240
ATCGTGTCTG	GCTTAAAGA	ATCAAAGAGC	GACAAAAGC	TGGCTCTCAA	GGAGCTCAAA	300
ACGAAGGGAG	ACACCTAAGA	AAATCTCTGA	TCAAGCTCAG	AGGAAGAAAA	TGTCTTAGAG	360
CTCAAGATGG	TATCCTTAAG	TACATTGTTG	AAGCTTATGG	AAGTCTGCAA	AGTTCGCGGG	420
TTTGTCTATG	GTATAATACC	GGAAAAGGGC	AAGCCTGTGA	GTGGGCTCCT	CTGACAATAT	480
AAGAGCTTGG	TGGAAAGAGA	AAGTGAAGTT	TGATAAGAAC	GGTCTGCTG	CTATTGCTAA	540
ATACGAAGAG	GAGTGTTTAG	CGTTTGGGAA	ATCTGATGGG	AATAGGAATT	CACAGTTTGT	600
TCTCCAGGAT	TTGCAAGATG	CTACTTTAGG	GTCTTTGTTA	TCTTCTTTGA	TGCAACATTG	660
TGATCCTCCT	CAAAGGAAGT	ATCCGTTGGA	GAAAGGGACG	CCTCCGCTT	GGTGGCCAAC	720
GGGGAATGAA	GAATGGTGGG	TGAAACTCGG	TCTGCCTAAA	AGCCAGAGTC	CTCCTTACCG	780
AAACCTCAT	GATCTCAAGA	AGATGTGGAA	GGTTGGAGTT	TTAACGGCAG	TGATCAATCA	840
TATGTTACCT	GATATTGCAA	AGATTAAAGAG	GCATGTTTCGT	CAGTCGAAAT	GTTTACAGGA	900
CAAGATACA	GCTAAAGAGA	GTGCGATTTG	GTTGGCGGTT	TTGAACCAAG	AGGAATCTTT	960
GATTCAGCAG	CCTAGCAGTG	ACAATGGAAA	CTCCAATGTG	ACTGAGACAC	ATCGTAGGGG	1020
TAATAACGCT	GACAGGAGGA	AACCTGTGGT	CAACAGTGAC	AGTGACTATG	ATGTTGATGG	1080
GACAGAGGAA	GCTTCAGGTT	CAGTTTCATC	TAAAGACAGT	AGAAGAAATC	AGATTCAAAA	1140
AGAACAAACCA	ACAGCCATCT	CACATTCACT	AAGAGATCAA	GATAAAGCAG	AGAAACATCG	1200

CAGAAGGAAA AGACCTCGAA TTAGATCCGG AACTGTCAAT CGACAAGAGG AAGAACAACC 1260
 TGAAGCTCAA CAAAGAAACA TCTTACCTGA TATGAATCAT GTTGATGCCC CTCTGCTAGA 1320
 ATATAACATC AACGGTACTC ATCAAGAGGA CGATGTTGTC GACCCAAATA TTGCCTTAGG 1380
 ACCAGAGGAT AATGGTCTGG AACTAGTGGT TCCTGAGTTC AATAACCAAA CATACTTATC 1440
 TTCCACTTGT TAATGAACAA ACTATGATGC CTGTAGACGA AAGGCCAATG CTTTATGGAC 1500
 CCAAACCCTA ACCAAGAGCT TCAATTTGGG TCAGGGTACA ACTTCTACAA TCCCTCTGCA 1560
 GTGTTTGTAC ATAACCAGGA AGACGACATT CTCCATACAC AGATAGAAAT GAATACACAA 1620
 GCACCACCTC ACAACAGTGG GTTCGAGGAG GCCCCAGGAG GAGTACTTCA ACCCCTTGGT 1680
 TTAICTCGGAA ATGAAGACGG TGTAACAGGG AGTGAGTTGC CTCAGTATCA GAGTGGCATT 1740
 CTGTCTCCAT TGA CTGACTT GGACTTTGAC TATGGTGGTT TTGGTGATGA TTTCTCATGG 1800
 TTTGGAGCTT AGTGTCTTGC CATT TTTT TTTT GGGAGATTAC ATAGTTCAAA AGGACATGGC 1860
 AATAGTCTGG CTAGTACAGT TACTTTCTCT TCTTCATTTC TTCTGATCTT ATATTCTTCC 1920
 TCTTTT TTTT TTATAATATT TTCTTAGATT TGTTAAGAGA AACAA TTTT CTTTGAATA 1980
 AGTTGCCAGA AGAACTGCTT TGCCCGTTGT AATGGTCTCT AGGGAAAGCA GTTAGCGTAT 2040
 CATCATTTGT AAATTTACCT GTGAG 2065

(2) INFORMATION FOR SEQ ID NO:12:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 567 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

Met Gly Asp Leu Ala Met Ser Val Ala Asp Ile Arg Met Glu Asn Glu
 1 5 10 15
 Pro Asp Asp Leu Ala Ser Asp Asn Val Ala Glu Ile Asp Val Ser Asp
 20 25 30
 Glu Glu Ile Asp Ala Asp Asp Leu Glu Arg Arg Met Trp Lys Asp Arg
 35 40 45
 Val Arg Leu Lys Arg Ile Lys Glu Arg Gln Lys Ala Gly Ser Gln Gly
 50 55 60
 Ala Gln Thr Lys Glu Thr Pro Lys Lys Ile Ser Asp Gln Ala Gln Arg
 65 70 75 80
 Lys Lys Met Ser Arg Ala Gln Asp Gly Ile Leu Lys Tyr Met Leu Lys
 85 90 95
 Leu Met Glu Val Cys Lys Val Arg Gly Phe Val Tyr Gly Ile Ile Pro

69

100										105					110					
Glu	Lys	Gly	Lys	Pro	Val	Ser	Gly	Ser	Ser	Asp	Asn	Ile	Arg	Ala	Trp					
		115					120					125								
Trp	Lys	Glu	Lys	Val	Lys	Phe	Asp	Lys	Asn	Gly	Pro	Ala	Ala	Ile	Ala					
	130					135					140									
Lys	Tyr	Glu	Glu	Glu	Cys	Leu	Ala	Phe	Gly	Lys	Ser	Asp	Gly	Asn	Arg					
145					150					155					160					
Asn	Ser	Gln	Phe	Val	Leu	Gln	Asp	Leu	Gln	Asp	Ala	Thr	Leu	Gly	Ser					
				165					170					175						
Leu	Leu	Ser	Ser	Leu	Met	Gln	His	Cys	Asp	Pro	Pro	Gln	Arg	Lys	Tyr					
			180					185					190							
Pro	Leu	Glu	Lys	Gly	Thr	Pro	Pro	Pro	Trp	Trp	Pro	Thr	Gly	Asn	Glu					
		195					200					205								
Glu	Trp	Trp	Val	Lys	Leu	Gly	Leu	Pro	Lys	Ser	Gln	Ser	Pro	Pro	Tyr					
	210					215					220									
Arg	Lys	Pro	His	Asp	Leu	Lys	Lys	Met	Trp	Lys	Val	Gly	Val	Leu	Thr					
225					230					235					240					
Ala	Val	Ile	Asn	His	Met	Leu	Pro	Asp	Ile	Ala	Lys	Ile	Lys	Arg	His					
				245					250					255						
Val	Arg	Gln	Ser	Lys	Cys	Leu	Gln	Asp	Lys	Met	Thr	Ala	Lys	Glu	Ser					
			260					265					270							
Ala	Ile	Trp	Leu	Ala	Val	Leu	Asn	Gln	Glu	Glu	Ser	Leu	Ile	Gln	Gln					
	275						280					285								
Pro	Ser	Ser	Asp	Asn	Gly	Asn	Ser	Asn	Val	Thr	Glu	Thr	His	Arg	Arg					
	290					295					300									
Gly	Asn	Asn	Ala	Asp	Arg	Arg	Lys	Pro	Val	Val	Asn	Ser	Asp	Ser	Asp					
305					310					315					320					
Tyr	Asp	Val	Asp	Gly	Thr	Glu	Glu	Ala	Ser	Gly	Ser	Val	Ser	Ser	Lys					
				325					330					335						
Asp	Ser	Arg	Arg	Asn	Gln	Ile	Gln	Lys	Glu	Gln	Pro	Thr	Ala	Ile	Ser					
			340					345					350							
His	Ser	Val	Arg	Asp	Gln	Asp	Lys	Ala	Glu	Lys	His	Arg	Arg	Arg	Lys					
	355						360					365								
Arg	Pro	Arg	Ile	Arg	Ser	Gly	Thr	Val	Asn	Arg	Gln	Glu	Glu	Glu	Gln					
	370					375					380									
Pro	Glu	Ala	Gln	Gln	Arg															

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Pro Asn Pro Asn Gln Glu Leu Gln Phe Gly Ser Gly Tyr Asn Phe Tyr
465                      470                      475                      480

Asn Pro Ser Ala Val Phe Val His Asn Gln Glu Asp Asp Ile Leu His
                      485                      490                      495

Thr Gln Ile Glu Met Asn Thr Gln Ala Pro Pro His Asn Ser Gly Phe
                    500                      505                      510

Glu Glu Ala Pro Gly Gly Val Leu Gln Pro Leu Gly Leu Leu Gly Asn
                    515                      520                      525

Glu Asp Gly Val Thr Gly Ser Glu Leu Pro Gln Tyr Gln Ser Gly Ile
530                      535                      540

Leu Ser Pro Leu Thr Asp Leu Asp Phe Asp Tyr Gly Gly Phe Gly Asp
545                      550                      555                      560

Asp Phe Ser Trp Phe Gly Ala
                    565

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(2) INFORMATION FOR SEQ ID NO:13:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

```

Met Thr Val Val Arg Glu Tyr Asp Pro Thr Arg Asp Leu Val Gly Val
1                      5                      10                      15

Glu Asp Val Glu Arg Arg Cys Glu Val Gly Pro Ser Gly Lys Leu Ser
20                      25                      30

Leu Phe Thr Asp Leu Leu Gly Asp Pro Ile Cys Arg Ile Arg His Ser
35                      40                      45

Pro Ser Tyr Leu Met Leu Val Ala Glu Met Gly Thr Glu Xaa Xaa Xaa
50                      55                      60

Lys Lys Glu Ile Val Gly Met Ile Arg Gly Cys Ile Lys Thr Val Thr
65                      70                      75                      80

Cys Gly Gln Lys Leu Asp Leu Asn His Lys Xaa Xaa Xaa Ser Gln Asn
85                      90                      95

Asp Val Val Xaa Xaa Lys Pro Leu Tyr Thr Lys Leu Xaa Xaa Xaa
100                      105                      110

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Ala Tyr Val Leu Gly Leu Arg Val
115                      120                      125

Ser Pro Phe His Arg Arg Gln Gly Ile Gly Phe Lys Leu Val Lys Met
130                      135                      140

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SUBSTITUTE SHEET (RULE 26)

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Met Glu Glu Trp Phe Arg Gln Xaa Asn Gly Ala Glu Tyr Ser Tyr Ile
 145 150 155 160

Ala Thr Glu Asn Asp Xaa Xaa Xaa Xaa Asn Gln Ala Ser Val Asn Leu
 165 170 175

Phe Thr Gly Lys Cys Gly Tyr Ser Glu Phe Arg Thr Pro Ser Ile Leu
 180 185 190

Val Asn Pro Val Tyr Ala His Arg Val Asn Val Ser Arg Arg Val Thr
 195 200 205

Val Ile Lys Leu Glu Pro Val Asp Ala Glu Thr Xaa Xaa Xaa Leu Tyr
 210 215 220

Arg Ile Arg Phe Ser Thr Thr Glu Phe Phe Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:14:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 1702 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:

CTCCAACTTT TAAAACATCAT CATAAATAGT AAAAAAGTAG CCGGAAAAAT AAAATAAAAA	60
GTCTATTTCT CTTTCCTTTA AAATCCAAAT CCTATAAACT CATAGCTTTC TCTGTTCTTT	120
ACTTATACCT CACGTTATAC ATATATATAG AGTTTCTATA AATGCTTCTC TTTCTCTCTG	180
AACAAATCTT CCTCACTTCT CTCATTTCCA CACTCACCTT CCTCTCTATA TATTAAACCC	240
TATCTACTTA ACTCTTCTTC TAACTCTAAT CTCTCTCTCT ATTTACTCTG CTTCTGTTCT	300
CACTCTGAAA GAACCAAAAC ATGACGGTGG TTAGAGAGTA CGACCCGACC CGAGACTTAG	360
TCGGCGTGGA GGACGTGGAA CGACGGTGTG AAGTCGGACC AAGCGGCAAG CTTTCTCTTT	420
TCACCGACCT TTTGGGTGAC CCGATTTGTA GAATCCGACA TTCACCTTCC TATCTCATGC	480
TGGTGGCTGA GATGGGTACG GAGAAGAAGG AGATAGTGGG CATGATTAGA GGATGTATCA	540
AAACCGTTAC ATGTGGCCAA AACTCGATT TAAATCACAA ATCTCAAAAC GATGTCGTTA	600
AGCCTCTTTA CACTAAACTC GCTTACGTCT TGGGCCTTCG CGTCTCTCCT TTTCACAGGA	660
GACAAGGGAT TGGGTTTAAG CTCGTGAAGA TGATGGAGGA ATGGTTTAGA CAAAACGGAG	720
CTGAGTATTC GTATATTGCA ACTGAGAACG ATAATCAAGC TTCTGTGAAT TTGTTACCCG	780
GGAAATGTGG TTATTCGGAG TTTCTGACAC CGTCGATTTT GGTAAACCCG GTTTACGCTC	840
ATCGAGTTAA TGTTTCGCGG CGAGTCACGG TTATCAAGTT AGAGCCGGTT GATGCTGAGA	900

SUBSTITUTE SHEET (RULE 26)

CGTTGTACCG AATCCGGTTT AGCACAACAG A3TTTTTCCC GCGGGATATT GATTCCGGTAC	960
TTAATAACAA ACTCTCGCTT GGGACTTTTCG TCGCGGTGCC ACGTGGAAGC TGTTATGGAT	1020
CCGGGTCTGG ATCATGGCCC GGTTCGGCTA AATTCTCGA ATATCCACCC GAGTCATGGG	1080
CCGTATTAAG CGTGTGGAAT TGTAAGACT CGTTTCTGTT AGAAGTACGT GGAGCGTCGA	1140
GATTGAGACG TGTGGTGGCT AAAACGACGC GAGTAGTTGA TAAAACGTTG CCGTTTCTGA	1200
AACTACCTTC GATACCGTCC GTTTTCGAAC CTTTGGACT TCATTTTATG TATGGAATCG	1260
GAGGAGAAGG TCCACGCGCG GTGAAGATGG TGAAATCCTT GTGTGCTCAC GCGCATAACT	1320
TGGCTAAGGC AGGTGGTTGT GGTGTCGTGG CGGCGGAAGT TGCCGGAGAA GACCCGTTGC	1380
GGCGAGGAAT ACCACATTGG AAAGTGCTAT CGTGTGACGA GGATCTTTGG TGTATAAAGC	1440
GGCTTGAGGA TGACTATAGT GATGGTGTG TTGGTGATTG GACTAAATCG CCACCTGGCG	1500
TTTCCATTTT TGTAAGCCCT AGAGAATTTT AAAACTTTTT TTTTAACTCT ATAATATATA	1560
TTCTCTATTA ACCACTTGAT GTTAAATTAG GGGTTTCTT CTAAGTTTAT AGATTTTCTT	1620
GTTTTAGAAT TAATCTTTTT TTTAGGTAAC TTTTTTGCT TTTGTTTTG TTTGTTTTG	1680
TTTTTGTGGG TGTATAAAT TA	1702

(2) INFORMATION FOR SEQ ID NO:15:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 4146 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:

TGTCATAATC AGTACAAAAT AAATCACCTA CCAACCTGAA CTATATGTTA TATATTTTGA	60
GGGGCCACGT CAAGTGTGCC GTTATTTTTT GTGTTTATGA TTGTTAATA TTTGTGCGTG	120
TGATGGTGTG TCTTGCTTAG TTTCCACTTA ATACACAATC AAATATCAAG TGGAATATT	180
TATGAAAATT GTTCTTCGAG AAGAATTCTG ACCCTAAAAG GTCATTTGAG GGCTTGAGGC	240
TTATTGTTTC CAAATTACAC CAGTAAACAA GGGTTTTTTT TTGTCAACAA AGATTATTGT	300
AATTCGAATT TCGTCTACAA TAAAACAATT TTCTTACTAA AACAAAACAA TTAGCTGACG	360
GTTGATATTT CGGCTTTTGA GTTTAATTAA CTAATTGGTG ATTATGTTGA TGATCTTTCA	420
CACCTAATGA AGTGTGATGT ATATGTATAT ATGTATATAC TTATGTATAT ATAAAACGTA	480
CATATAATCA TTTGTCATAT ATATCATCAT GTATTGCATG ACTAAACTAC CCTTAAAAGA	540
GGAATACGAT AGACATGACC TTTAGGAATT TGTTTTTTTC TTCTAAATGG ATTCCTTCGC	600
TTCTTTTTAG CCTCGTAGTG AATTGAACA TTGCAGTTAT TTCTAGTAAG ATATTTTTTC	660

TGTATTTTTC	GGAAATGTT	AAAACTAAT	TATACACAAT	TTACTTTCTC	TCTCAACTCT	720
TATTTTACGT	TACTGTTTTT	TTTTTCCTCT	TGCAAAATTA	GAGCTGATGT	ATTTACATTT	780
ACTAGTAATT	TGGTAGATAG	ACAGTTAATG	TAGTATATAG	ATGGGGTTGA	GGGCAAATGA	840
TTACTTGGGA	GATGGTGCAA	TGCATCAGAG	TGATGATGTG	GAATTTAATA	AGTGTGAATT	900
TATGGGCAAA	GGAAGGGAAC	TAGTAGTAGA	AAGGGAAATA	AATACAGTAC	AAGTAAGAGG	960
AAAACGAAAA	GAGAGATAGA	AACCATAATA	ATGAGTTAAC	GCAGACATAG	CCGCCATTTT	1020
CAACTTCTCA	CTCCCACTTA	CAACTTCTCC	ITCTGGGCAA	GTTTTCCACA	TCAATGCTCG	1080
TCTTAATCAC	CATTAATCTC	TACTCATCAT	TAATACGTTG	AAGCCCACTA	TTTCAAAATT	1140
TACTAGGAGT	ATTTATTCGT	GAAAAACATT	TAAATGTCCC	TAATTATAAG	AGATTTAATT	1200
TCATATTTAT	TGTATTAAAG	AGAATTTACA	TTAGCTGTCA	AAAAAAAAAA	AAAAAGAGAA	1260
TTAACATTAT	TTTACAGAAC	ATAAAATTTT	GAAAAATAGAT	AGCGCCACTG	CATGTAAGAA	1320
CATACAAATT	TCTTTTTTTC	AACAAAATCT	ATTATATTTT	CTTCTTTTTT	TGAACATTAT	1380
GTGTAGTTTG	TAGTAACTA	AAAAGTGTGG	ACCAACACAA	TTTAAATCAT	TCGATTTTGT	1440
AGCAAAAACA	TTTTTGTTCC	AATTTCCAAG	CAGCAAATAT	GGAAGGAATA	TAAATTCCTT	1500
ACTATTTTTTC	CTCTTAACAC	ATAAAAGTAA	AAAAAGCATT	CAATGATCAG	TTAAATCTG	1560
GTTAGAATTC	TACCTTATCA	TTTAGAACTA	GCTAATATTT	AAATTCATAT	ATACAAAAAA	1620
TAAAATGGGA	ACTGTAGAGA	CTAGAGACTA	TAAATAGAGG	ATTGAGAAGA	AGAACTTTTA	1680
AAGCTCTATC	AATCATGAAC	TACTCGCCTT	CTCCAACCTT	TAAAACTCAT	CATAAATAGT	1740
AAAAAAGTAG	CCGGAAAAAT	AAAATAAAAA	GTCTATTTCT	CTTTCCTTTA	AAATCCAAAT	1800
CCTATAAACT	CATAGCTTTC	TCTGTTCTTT	ACTTATACCT	CACGTTATAC	ATATATATAG	1860
AGTTTCTATA	AATGCTTCTC	TTTCCTCTCG	AACAAATCTT	CCTCACTTCT	CTCATTTCCA	1920
CACTCACCTT	CCTCTCTATA	TATTAAACCC	TATCTACTTA	ACTCTTCTTC	TAACTCTAAT	1980
CTCTCTCTCT	ATTTACTCTG	CTTCTGTTCT	CACTCTGAAA	GAACCAAAAC	ATGACGGTGG	2040
TTAGAGAGTA	CGACCCGACC	CGAGACTTAG	TCGGCGTGGA	GGACGTGGAA	CGACGGTGTG	2100
AAGTCGGACC	AAGCGGCAAG	CTTCTCTTTT	TCACCGACCT	TTTGGGTGAC	CCGATTTGTA	2160
GAATCCGACA	TTCACCTTCC	TATCTCATGC	TGGTAATAAC	ATGTTTCACA	ATCTTTTATC	2220
TTCTTTTACT	TGTATGTCTC	TTCAAAAAT	CTGTTTGTTT	TTTGAACCTA	GAAGTAGAAA	2280
ACATAGAACA	CCAACCTTCT	AACCTTTGGT	TAATCCAAAA	AACCCATTTT	CCATAAACAA	2340
TTAAAGTTCG	GTTCTTTTTT	TGGTATCATT	TCTATTTTTT	TCCGATTCTT	GATAAGATCA	2400
AAAGACTCAT	CATTTATATT	ATTTTTTGCA	ACCAAATGAT	ACCCGAGTAA	CTATAACTAA	2460
TAAAGTTTCC	TCTTTATTAT	AAAAGGTTAA	AAACATATAA	TAACGGAAAA	TTTAAATTAT	2520
GGGACTGTAA	CAGGTGGCTG	AGATGGGTAC	GGAGAAGAAG	GAGATAGTGG	GCATGATTAG	2580
AGGATGTATC	AAAACCGTTA	CATGTGGCCA	AAAACCTCGAT	TTAAATCACA	AATCTCAAAA	2640
CGATGTCGTT	AAGCCTCTTT	ACACTAAACT	CGCTTACGTC	TTGGGCCTTC	GCGTCTCTCC	2700

TTTTCACAGG	TACCCCTTCCG	TTTTCCTCCC	ACTCATAATC	ACACGCTATT	ATAGATTTTG	2760
GTTATCTAAA	CTAGTTTTGG	TTTTTGCAGG	AGACAAGGGA	TGGGTTTAA	GCTCGTGAAG	2820
ATGATGGAGG	AATGGTTTAG	ACAAAACGGA	GCTGAGTATT	CGTATATTGC	AACTGAGAAC	2880
GATAATCAAG	CTTCTGTGAA	TTTGTTCAAC	GGGAAATGTG	GTTATTTCGGA	GTTTCGTACA	2940
CCGTCGATTT	TGGTTAACCC	GGTTTACGCT	CATCGAGTTA	ATGTTTCGCG	GCGAGTCACG	3000
GTTATCAAGT	TAGAGCCGGT	TGATGCTGAG	ACGTGTACC	GAATCCGGT	TAGCACAACA	3060
GAGTTTTTCC	CGCGGGATAT	TGATTTCGTA	CTTAATAACA	AACTCTCGCT	TGGGACTTTC	3120
GTCGCGGTGC	CACGTGGAAG	CTGTTATGGA	TCCGGGTCTG	GATCATGGCC	CGGTTCCGGCT	3180
AAATTCCTCG	AATATCCACC	CGAGTCATGG	GCCGTATTAA	GCGTGTGGAA	TTGTAAAGAC	3240
TCGTTTCTGT	TAGAAGTACG	TGGAGCGTCG	AGATTGAGAC	GTGTGGTGGC	TAAAACGACG	3300
CGAGTAGTTG	ATAAAACGTT	GCCGTTTCTG	AACTACCTT	CGATACCGTC	CGTTTTTCGAA	3360
CCTTTTGGAC	TTCATTTTAT	GTATGGAATC	GGAGGAGAAG	GTCCACGCGC	GGTGAAGATG	3420
GTGAAATCCT	TGTGTGCTCA	CGCGCATAAC	TTGGCTAAGG	CAGGTGGTTG	TGGTGTCTGT	3480
GCGGCGGAAG	TTGCCGGAGA	AGACCCGTTG	CGGCGAGGAA	TACCACATTG	GAAAGTGCTA	3540
TCGTGTGACG	AGGATCTTTG	GTGTATAAAG	CGGCTTGGAG	ATGACTATAG	TGATGGTGTT	3600
GTTGGTGATT	GGACTAAATC	GCCACCTGGC	GTTTCCATTT	TTGTAGACCC	TAGAGAATTT	3660
TAAAACTTTT	TTTTTAACCT	TATAATATAT	ATTCTCTATT	AACCACTTGA	TGTTAAATTA	3720
GGGGTTTTCT	TCTAAGTTTA	TAGATTTTCT	TGTTTTAGAA	TTAATCTTTT	TTTTAGGTAA	3780
CTTTTTTTGC	TTTTTGTTTT	GTTTTGTTTT	GTTTTTGTGG	GTGTTATAAA	TTAGTGGTAA	3840
GAGGTAATAT	CTCCTACTTT	TGGGTTTGTG	TCTTCTTGTC	TTGTAAATGG	ATCTAGCTTT	3900
TTAAGATACT	TTTTCTTTGT	GGCCAAACCA	AAACGCCGAC	CTGATTATTA	TTTCCAAGTA	3960
GATAAAATTT	CATGAACGCA	CTGATACGTA	TAATGATGCA	ATTTGTGTTA	AGACGATACT	4020
TTGGAGATAA	AATTACAATA	TGACAATGAT	AGAAAATGTT	ACCAATAACG	ATTAGCATT	4080
TCGTGTGTGC	CATCAAGTAT	AACTAAGAGA	AAGACGCACA	TTTTCTTTAA	GAGTAAATAA	4140
AATATT						4146

(2) INFORMATION FOR SEQ ID NO:16:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 398 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

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Met Thr Val Val Arg Glu Tyr Asp Pro Thr Arg Asp Leu Val Gly Val
 1 5 10 15
 Glu Asp Val Glu Arg Arg Cys Glu Val Gly Pro Ser Gly Lys Leu Ser
 20 25 30
 Leu Phe Thr Asp Leu Leu Gly Asp Pro Ile Cys Arg Ile Arg His Ser
 35 40 45
 Pro Ser Tyr Leu Met Leu Val Ala Glu Met Gly Thr Glu Lys Lys Glu
 50 55 60
 Ile Val Gly Met Ile Arg Gly Cys Ile Lys Thr Val Thr Cys Gly Gln
 65 70 75 80
 Lys Leu Asp Leu Asn His Lys Ser Gln Asn Asp Val Val Lys Pro Leu
 85 90 95
 Tyr Thr Lys Leu Ala Tyr Val Leu Gly Leu Arg Val Ser Pro Phe His
 100 105 110
 Arg Arg Gln Gly Ile Gly Phe Lys Leu Val Lys Met Met Glu Glu Trp
 115 120 125
 Phe Arg Gln Asn Gly Ala Glu Tyr Ser Tyr Ile Ala Thr Glu Asn Asp
 130 135 140
 Asn Gln Ala Ser Val Asn Leu Phe Thr Gly Lys Cys Gly Tyr Ser Glu
 145 150 155 160
 Phe Arg Thr Pro Ser Ile Leu Val Asn Pro Val Tyr Ala His Arg Val
 165 170 175
 Asn Val Ser Arg Arg Val Thr Val Ile Lys Leu Glu Pro Val Asp Ala
 180 185 190
 Glu Thr Leu Tyr Arg Ile Arg Phe Ser Thr Thr Glu Phe Phe Pro Arg
 195 200 205
 Asp Ile Asp Ser Val Leu Asn Asn Lys Leu Ser Leu Gly Thr Phe Val
 210 215 220
 Ala Val Pro Arg Gly Ser Cys Tyr Gly Ser Gly Ser Gly Ser Trp Pro
 225 230 235 240
 Gly Ser Ala Lys Phe Leu Glu Tyr Pro Pro Glu Ser Trp Ala Val Leu
 245 250 255
 Ser Val Trp Asn Cys Lys Asp Ser Phe Leu Leu Glu Val Arg Gly Ala
 260 265 270
 Ser Arg Leu Arg Arg Val Val Ala Lys Thr Arg Arg Val Val Asp Lys
 275 280 285
 Thr Leu Pro Phe Leu Lys Leu Pro Ser Ile Pro Ser Val Phe Glu Pro
 290 295 300
 Phe Gly Leu His Phe Met Tyr Gly Ile Gly Gly Glu Gly Pro Arg Ala
 305 310 315 320
 Val Lys Met Val Lys Ser Leu Cys Ala His Ala His Asn Leu Ala Lys
 325 330 335
 Ala Gly Gly Cys Gly Val Val Ala Ala Glu Val Ala Gly Glu Asp Pro
 340 345 350
 Leu Arg Arg Gly Ile Pro His Trp Lys Val Leu Ser Cys Asp Glu Asp

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355		360		365
Leu Trp Cys Ile Lys Arg	Leu Gly Asp Asp Tyr Ser Asp Gly Val Val			
370	375		380	
Gly Asp Trp Thr Lys Cys His Leu Ala Phe Pro Phe Leu Glx				
385	390		395	

(2) INFORMATION FOR SEQ ID NO:17:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 12 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

GAGTTGCGCA TG

12

(2) INFORMATION FOR SEQ ID NO:18:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 4 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

Gly Val Ala His
1

(2) INFORMATION FOR SEQ ID NO:19:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:19:

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TGCTACAATC AGAATTCTTG CAGT

24

(2) INFORMATION FOR SEQ ID NO:20:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 8 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:

Ala Thr Ile Arg Ile Leu Ala Val
1 5

(2) INFORMATION FOR SEQ ID NO:21:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 23 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:

GGATCCTCTA GTCAATTAC CGC

23

(2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

AGATCTGGTA TATCCGTCT GCAC

24

(2) INFORMATION FOR SEQ ID NO:23:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid

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- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

CCGGATTTCGG TTTGTAGC

18

(2) INFORMATION FOR SEQ ID NO:24:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:

GACGTGCATG TTCTTGGG

18

(2) INFORMATION FOR SEQ ID NO:25:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:

GAAAGCCACA TCACCTGC

18

(2) INFORMATION FOR SEQ ID NO:26:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 17 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

79

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:

GGGGTGGAGT TATCCAC

17

(2) INFORMATION FOR SEQ ID NO:27:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 17 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:

GACACCGGGA AGTATCG

17

(2) INFORMATION FOR SEQ ID NO:28:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 19 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:28:

CTGCTTTCAT AGAAGAGGC

19

(2) INFORMATION FOR SEQ ID NO:29:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 19 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:29:

GTCAGAACAA ACCTGCTCC

19

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(2) INFORMATION FOR SEQ ID NO:30:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 17 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:

CACCCAGGTC TTGGTGG

17

(2) INFORMATION FOR SEQ ID NO:31:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 16 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:31:

GGCCGCCATG GATGCG

16

(2) INFORMATION FOR SEQ ID NO:32:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:

TCTCAATCAA GAGGAGGC

18

(2) INFORMATION FOR SEQ ID NO:33:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

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- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:

CTTGAAGGAT CCGAGTGG

18

(2) INFORMATION FOR SEQ ID NO:34:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:

CAGGTTGGCG AGTTCCTCG

19

(2) INFORMATION FOR SEQ ID NO:35:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:

CTTGCTGTGA TTCTCCATGC

20

(2) INFORMATION FOR SEQ ID NO:36:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:

CCCTGGACCA GCTCCTGG

18

(2) INFORMATION FOR SEQ ID NO:37:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

TGGCGCAAGC ATCGTCCC

18

(2) INFORMATION FOR SEQ ID NO:38:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:38:

AAATGTTTCAG GAATCTCTCG

20

(2) INFORMATION FOR SEQ ID NO:39:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:39:

CTGGCTGGCA GCCACGCC

18

(2) INFORMATION FOR SEQ ID NO:40:

- (i) SEQUENCE CHARACTERISTICS:

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- (A) LENGTH: 18 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:

GCGTTCTCAA AGCTGCGG

18

(2) INFORMATION FOR SEQ ID NO:41:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:

ACTGATGGGT CTTCTGGG

18

(2) INFORMATION FOR SEQ ID NO:42:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:42:

GGATCAGGAT GGACCCGG

18

(2) INFORMATION FOR SEQ ID NO:43:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

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(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:

TGGTTGCTGA AGCCAGGG

18

(2) INFORMATION FOR SEQ ID NO:44:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:

TCCATTCATA GAGAGTGGG

19

(2) INFORMATION FOR SEQ ID NO:45:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:45:

ATGCCCAAGA ACATGCACG

19

(2) INFORMATION FOR SEQ ID NO:46:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:

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CAACTGATCC TTTACCCTGC

(2) INFORMATION FOR SEQ ID NO:47:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:47:

19

GTTGTTAGGT CAACTTGCG

(2) INFORMATION FOR SEQ ID NO:48:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:48:

19

CTCTGTTAGG GCTTCCTCC

(2) INFORMATION FOR SEQ ID NO:49:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:49:

18

GAATCAGATT TCGCGAGG

(2) INFORMATION FOR SEQ ID NO:50:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid

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(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO
(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:50:

GTCCAAATGG AGGAAGCC

18

(2) INFORMATION FOR SEQ ID NO:51:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 23 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO
(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:51:

CCACGACTGT ACAATTGACC TTG

23

(2) INFORMATION FOR SEQ ID NO:52:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 18 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO
(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:

CATGATCGCA AGTTGACC

18

(2) INFORMATION FOR SEQ ID NO:53:

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 22 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA
(iii) HYPOTHETICAL: NO

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(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:

AGAAAACCTCT TATCAAGCTA CG

22

(2) INFORMATION FOR SEQ ID NO:54:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 20 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:

AAGCTTATGG GTGCTCGTGC

20

(2) INFORMATION FOR SEQ ID NO:55:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 20 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:55:

GGAAAGAGAG AAAGACTCAG

20

(2) INFORMATION FOR SEQ ID NO:56:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 18 base pairs
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:56:

GCCACCAAGT CATACCCG

18

88

(2) INFORMATION FOR SEQ ID NO:57:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:57:

CCTTCTATAT TTGGTTCC

18

(2) INFORMATION FOR SEQ ID NO:58:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:58:

CCATTCTCCG GAATAATCC

19

(2) INFORMATION FOR SEQ ID NO:59:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:59:

CACGGAGCAG GATAAGGGTA

20

(2) INFORMATION FOR SEQ ID NO:60:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

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- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:60:

CGGATTGGAT TGTGTGTGC

19

(2) INFORMATION FOR SEQ ID NO:61:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:

CGCCACTGCA TGTAAGAAC

19

(2) INFORMATION FOR SEQ ID NO:62:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:

TCCACACGCT TAATACGGC

19

(2) INFORMATION FOR SEQ ID NO:63:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: YES

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:63:

GGTACGGAGA AGAAGGAG

18

(2) INFORMATION FOR SEQ ID NO:64:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:64:

CGCGGGATAT TGATTCGGT

19

(2) INFORMATION FOR SEQ ID NO:65:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 19 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:65:

GTGTTGAACA CGCCACAA

19

(2) INFORMATION FOR SEQ ID NO:66:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:66:

ACGACACCAC AACCACCT

18

(2) INFORMATION FOR SEQ ID NO:67:

- (i) SEQUENCE CHARACTERISTICS:

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- (A) LENGTH: 18 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:67:

GACAAGAAGA CACAAACC

18

(2) INFORMATION FOR SEQ ID NO:68:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: YES

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:68:

GAATCGGAGG AGAAGGTC

18

(2) INFORMATION FOR SEQ ID NO:69:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 240 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:69:

Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa
1				5					10							15
Xaa	Met	Phe	Gly	Tyr	Arg	Ser	Asn	Val	Pro	Lys	Val	Arg	Leu	Thr	Thr	
			20					25					30			
Asp	Arg	Leu	Val	Val	Arg	Leu	Val	His	Asp	Arg	Asp	Ala	Trp	Arg	Leu	
		35					40					45				
Ala	Asp	Tyr	Tyr	Ala	Glu	Asn	Arg	His	Phe	Leu	Lys	Pro	Trp	Glu	Pro	
		50				55					60					

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Val Arg Asp Glu Ser His Cys Tyr Pro Ser Gly Trp Gln Ala Arg Leu
65              70              75              80

Gly Met Ile Ash Glu Phe His Lys Gln Gly Ser Ala Phe Tyr Phe Gly
85              90              95

Leu Phe Asp Pro Asp Glu Lys Glu Ile Ile Gly Val Ala Asn Phe Ser
100            105            110

Asn Val Val Arg Gly Ser Phe His Ala Cys Tyr Leu Gly Tyr Ser Ile
115            120            125

Gly Gln Lys Trp Gln Gly Lys Gly Leu Met Phe Glu Ala Leu Thr Ala
130            135            140

Ala Ile Arg Tyr Met Gln Arg Thr Gln His Ile His Arg Ile Met Ala
145            150            155            160

Asn Tyr Met Pro His Xaa Xaa Xaa Xaa Asn Lys Arg Ser Gly Asp Leu
165            170            175

Leu Ala Arg Leu Gly Phe Glu Lys Glu Gly Tyr Ala Lys Asp Tyr Leu
180            185            190

Leu Ile Asp Gly Gln Trp Arg Asp His Val Leu Thr Ala Leu Thr Thr
195            200            205

Pro Asp Trp Thr Pro Gly Arg Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
210            215            220

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
225            230            235            240

```

(2) INFORMATION FOR SEQ ID NO:70:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 240 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:70:

```

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
1              5              10              15

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met Glu Thr Glu Ile Lys Val Ser
20            25            30

Glu Ser Leu Glu Leu His Ala Val Ala Glu Asn His Val Lys Pro Leu
35            40            45

Tyr Gln Leu Ile Cys Lys Asn Lys Thr Trp Leu Gln Gln Ser Leu Asn
50            55            60

Trp Pro Gln Phe Val Gln Ser Glu Glu Asp Thr Arg Lys Thr Val Gln
65            70            75            80

```

SUBSTITUTE SHEET (RULE 26)

93

Gly Asn Val Xaa Met Leu His Gln Arg Gly Tyr Ala Lys Met Phe Met
 85 90 95
 Ile Phe Xaa Xaa Lys Glu Asp Glu Leu Ile Gly Val Ile Ser Phe Xaa
 100 105 110
 Asn Arg Ile Glu Pro Leu Asn Lys Thr Ala Glu Ile Gly Tyr Trp Leu
 115 120 125
 Asp Glu Ser His Gln Gly Gln Gly Ile Ile Ser Gln Ala Leu Gln Ala
 130 135 140
 Leu Ile His His Tyr Ala Gln Ser Gly Glu Leu Arg Arg Phe Val Ile
 145 150 155 160
 Lys Cys Arg Val Asp Xaa Xaa Xaa Xaa Asn Pro Gln Ser Asn Gln Val
 165 170 175
 Ala Leu Arg Asn Gly Phe Ile Leu Glu Gly Cys Leu Lys Gln Ala Glu
 180 185 190
 Phe Leu Asn Asp Ala Tyr Asp Asp Val Asn Leu Tyr Ala Arg Ile Ile
 195 200 205
 Asp Ser Gln Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 210 215 220
 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:71:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:71:

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met Leu Trp Ser Ser Asn Asp Val Thr
 1 5 10 15
 Gln Gln Gly Ser Arg Pro Lys Thr Lys Leu Gly Gly Ser Xaa Met Ser
 20 25 30
 Ile Ile Ala Thr Val Lys Ile Gly Pro Asp Glu Ile Ser Ala Met Arg
 35 40 45
 Ala Val Leu Asp Leu Phe Gly Lys Glu Phe Glu Asp Ile Pro Thr Tyr
 50 55 60
 Ser Asp Arg Gln Pro Thr Asn Glu Tyr Leu Ala Asn Leu Leu His Ser
 65 70 75 80
 Glu Thr Phe Ile Ala Leu Ala Ala Phe Asp Arg Gly Thr Ala Ile Gly
 85 90 95

94

```

Gly Leu Ala Xaa Xaa Ala Tyr Val Leu Pro Lys Phe Glu Gln Ala Arg
      100      105      110
Ser Glu Xaa Xaa Xaa Xaa Xaa Ile Tyr Ile Tyr Asp Leu Ala Val
      115      120      125
Ala Ser Ser His Arg Arg Leu Gly Val Ala Thr Ala Leu Ile Ser His
      130      135      140
Leu Lys Arg Xaa Val Ala Val Glu Leu Gly Ala Tyr Val Ile Tyr Val
      145      150      155      160
Gln Ala Asp Tyr Gly Xaa Xaa Xaa Xaa Asp Asp Pro Ala Val Ala Leu
      165      170      175
Tyr Thr Lys Leu Gly Val Arg Glu Asp Val Met His Phe Asp Ile Asp
      180      185      190
Pro Arg Thr Ala Thr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
      195      200      205
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
      210      215      220
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
      225      230      235      240

```

(2) INFORMATION FOR SEQ ID NO:72:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:72:

```

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met Leu Arg Ser Ser Asn Asp Val Thr
1      5      10      15
Gln Gln Gly Ser Arg Pro Lys Thr Lys Leu Gly Gly Ser Ser Met Gly
      20      25      30
Ile Ile Arg Thr Cys Arg Leu Gly Pro Asp Gln Val Lys Ser Met Arg
      35      40      45
Ala Ala Leu Asp Leu Phe Gly Arg Glu Phe Gly Asp Val Ala Thr Tyr
      50      55      60
Ser Gln His Gln Pro Asp Ser Asp Tyr Leu Gly Asn Leu Leu Arg Ser
      65      70      75      80
Lys Thr Phe Ile Ala Leu Ala Ala Phe Asp Gln Glu Ala Val Val Gly
      85      90      95
Ala Leu Ala Xaa Xaa Ala Tyr Val Leu Pro Lys Phe Glu Gln Ala Arg
      100      105      110

```

SUBSTITUTE SHEET (RULE 26)

95

```

Ser Glu Xaa Xaa Xaa Xaa Xaa Xaa Ile Tyr Ile Tyr Asp Leu Ala Val
   115                               120                               125

Ser Gly Glu His Arg Arg Gln Gly Ile Ala Thr Ala Leu Ile Asn Leu
   130                               135                               140

Leu Lys His Xaa Glu Ala Asn Ala Leu Gly Ala Tyr Val Ile Tyr Val
   145                               150                               155                               160

Gln Ala Asp Tyr Gly Xaa Xaa Xaa Xaa Asp Asp Pro Ala Val Ala Leu
                   165                               170                               175

Tyr Thr Lys Leu Gly Ile Arg Glu Glu Val Met His Phe Asp Ile Asp
                   180                               185                               190

Pro Ser Thr Ala Thr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
   195                               200                               205

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
   210                               215                               220

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
   225                               230                               235                               240

```

(2) INFORMATION FOR SEQ ID NO:73:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:73:

```

Met Thr Thr Leu Asp Asp Thr Ala Tyr Arg Tyr Arg Thr Ser Val Pro
 1                               5                               10                               15

Gly Asp Ala Glu Ala Ile Glu Ala Leu Asp Gly Ser Phe Thr Thr Asp
                   20                               25                               30

Thr Val Phe Arg Val Thr Ala Thr Gly Asp Gly Phe Thr Leu Arg Glu
   35                               40                               45

Val Pro Val Asp Pro Pro Leu Thr Lys Val Xaa Xaa Phe Pro Asp Asp
   50                               55                               60

Glu Ser Asp Asp Glu Ser Asp Asp Gly Glu Asp Gly Asp Pro Asp Ser
   65                               70                               75                               80

Arg Thr Phe Val Ala Tyr Gly Asp Xaa Xaa Xaa Xaa Xaa Xaa Asp Gly
                   85                               90                               95

Asp Leu Ala Xaa Xaa Gly Phe Val Val Ile Ser Tyr Ser Ala Trp Asn
   100                               105                               110

Arg Arg Xaa Xaa Xaa Xaa Xaa Xaa Leu Thr Val Glu Asp Ile Glu Val
   115                               120                               125

```


96

Ala Pro Glu His Arg Gly His Gly Val Gly Arg Ala Leu Met Gly Leu
 130 135 140

Ala Thr Glu Xaa Phe Ala Gly Glu Arg Gly Ala Gly His Leu Trp Leu
 145 150 155 160

Glu Val Thr Asn Val Xaa Xaa Xaa Xaa Asn Ala Pro Ala Ile His Ala
 165 170 175

Tyr Arg Arg Met Gly Phe Thr Leu Cys Gly Leu Asp Thr Ala Leu Tyr
 180 185 190

Asp Gly Thr Ala Ser Asp Gly Glu Arg Gln Ala Leu Tyr Met Ser Met
 195 200 205

Pro Cys Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 210 215 220

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:74:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:74:

Met Thr Thr Thr His Gly Ser Thr Tyr Glu Phe Arg Ser Ala Arg Pro
 1 5 10 15

Gly Asp Ala Glu Ala Ile Glu Gly Leu Asp Gly Ser Phe Thr Thr Ser
 20 25 30

Thr Val Phe Glu Val Asp Val Thr Gly Asp Gly Phe Ala Leu Arg Glu
 35 40 45

Val Pro Ala Asp Pro Pro Leu Val Lys Val Xaa Xaa Phe Pro Asp Asp
 50 55 60

Gly Gly Ser Asp Gly Glu Asp Gly Ala Glu Gly Glu Asp Ala Asp Ser
 65 70 75 80

Arg Thr Phe Val Ala Val Gly Ala Xaa Xaa Xaa Xaa Xaa Xaa Asp Gly
 85 90 95

Asp Leu Ala Xaa Xaa Gly Phe Ala Ala Val Ser Tyr Ser Ala Trp Asn
 100 105 110

Gln Arg Xaa Xaa Xaa Xaa Xaa Xaa Leu Thr Ile Glu Asp Ile Glu Val
 115 120 125

Ala Pro Gly His Arg Gly Lys Gly Il Gly Arg Val Leu Met Arg His
 130 135 140

97

```

Ala Ala Asp Xaa Phe Ala Arg Glu Arg Gly Ala Gly His Leu Trp Leu
145          150          155
Glu Asn Thr Asn Val Xaa Xaa Xaa Xaa Asn Ala Pro Ala Ile His Ala
          165          170          175
Tyr Arg Arg Met Gly Phe Ala Phe Cys Gly Leu Asp Ser Ala Leu Tyr
          180          185          190
Gln Gly Thr Ala Ser Glu Gly Glu Xaa His Ala Leu Tyr Met Ser Met
          195          200          205
Pro Cys Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
210          215          220
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
225          230          235          240

```

(2) INFORMATION FOR SEQ ID NO:75:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:75:

```

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met Lys Ile Ser Val Ile Pro Glu
1          5          10
Gln Val Ala Glu Thr Leu Asp Ala Xaa Glu Asn His Phe Ile Val Arg
          20          25          30
Glu Val Phe Asp Val His Leu Ser Asp Gln Gly Phe Glu Leu Ser Thr
          35          40          45
Arg Ser Val Ser Pro Tyr Arg Lys Asp Tyr Xaa Xaa Ile Ser Asp Asp
          50          55          60
Asp Ser Asp Glu Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Asp Ser
65          70          75          80
Ala Cys Tyr Gly Ala Phe Xaa Ile Xaa Xaa Xaa Xaa Xaa Xaa Asp Gln
          85          90          95
Glu Leu Val Xaa Xaa Gly Lys Ile Glu Leu Asn Xaa Ser Thr Trp Asn
          100          105          110
Asp Leu Xaa Xaa Xaa Xaa Xaa Xaa Ala Ser Ile Glu His Ile Val Val
          115          120          125
Ser His Thr His Arg Gly Lys Gly Val Ala His Ser Leu Ile Glu Phe
          130          135          140
Ala Lys Lys Xaa Trp Ala Leu Ser Arg Gln Leu Leu Gly Ile Arg Leu
145          150          155          160

```

SUBSTITUTE SHEET (RULE 26)

98

Glu	Thr	Gln	Thr	Asn	Xaa	Xaa	Xaa	Xaa	Asn	Val	Pro	Ala	Cys	Asn	Leu
				165					170					175	
Tyr	Ala	Lys	Cys	Gly	Phe	Thr	Leu	Gly	Gly	Ile	Asp	Leu	Phe	Thr	Tyr
			180					185					190		
Lys	Thr	Arg	Pro	Gln	Val	Ser	Asn	Glu	Thr	Ala	Met	Tyr	Trp	Tyr	Trp
		195					200					205			
Phe	Ser	Gly	Ala	Gln	Asp	Asp	Ala	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa
	210					215						220			
Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa
225					230					235					240

(2) INFORMATION FOR SEQ ID NO:76:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 240 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:76:

Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Met
1				5					10						15
Ala	Lys	Phe	Lys	Ile	Arg	Pro	Ala	Thr	Ala	Ser	Asp	Cys	Ser	Xaa	Xaa
			20					25					30		
Xaa	Xaa	Asp	Ile	Leu	Arg	Leu	Ile	Lys	Glu	Leu	Ala	Lys	Tyr	Glu	Tyr
		35					40					45			
Met	Glu	Asp	Gln	Val	Ile	Leu	Thr	Glu	Lys	Asp	Leu	Gln	Glu	Asp	Gly
	50					55					60				
Phe	Gly	Glu	His	Pro	Phe	Tyr	His	Cys	Leu	Val	Ala	Glu	Val	Pro	Lys
65					70				75					80	
Glu	His	Trp	Thr	Pro	Xaa	Xaa	Xaa	Xaa	Xaa	Glu	Gly	His	Ser	Ile	Val
				85					90					95	
Gly	Phe	Ala	Xaa	Xaa	Met	Tyr	Tyr	Phe	Thr	Tyr	Asp	Pro	Trp	Ile	Gly
			100					105					110		
Lys	Leu	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Leu	Tyr	Leu	Glu	Asp	Phe	Phe	Val
	115						120					125			
Met	Ser	Asp	Tyr	Arg	Gly	Phe	Gly	Ile	Gly	Ser	Glu	Ile	Leu	Lys	Asn
	130					135					140				
Leu	Ser	Gln	Xaa	Val	Ala	Met	Lys	Cys	Arg	Cys	Ser	Ser	Met	His	Phe
145					150					155					160
Leu	Val	Ala	Glu	Trp	Xaa	Xaa	Xaa	Xaa	Asn	Glu	Pro	Ser	Ile	Asn	Phe
				165					170					175	

SUBSTITUTE SHEET (RULE 26)

99

```

Tyr Lys Arg Arg Gly Ala Ser Asp Leu Ser Ser Glu Glu Gly Trp Xaa
      180      185      190
Xaa Xaa Xaa Xaa Arg Leu Phe Lys Ile Asp Lys Glu Tyr Leu Leu Lys
      195      200      205
Met Ala Ala Glu Glu Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
      210      215      220
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
      225      230      235      240

```

(2) INFORMATION FOR SEQ ID NO:77:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 240 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:77:

```

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met
 1      5      10      15
Ala Lys Phe Val Ile Arg Pro Ala Thr Ala Ala Asp Cys Ser Xaa Xaa
 20      25      30
Xaa Xaa Asp Ile Leu Arg Leu Ile Lys Glu Leu Ala Lys Tyr Glu Tyr
 35      40      45
Met Glu Glu Gln Val Ile Leu Thr Glu Lys Asp Leu Leu Glu Asp Gly
 50      55      60
Phe Gly Glu His Pro Phe Tyr His Cys Leu Val Ala Glu Val Pro Lys
 65      70      75      80
Glu His Trp Thr Pro Xaa Xaa Xaa Xaa Xaa Glu Gly His Ser Ile Val
 85      90      95
Gly Phe Ala Xaa Xaa Met Tyr Tyr Phe Thr Tyr Asp Pro Trp Ile Gly
100      105      110
Lys Leu Xaa Xaa Xaa Xaa Xaa Xaa Leu Tyr Leu Glu Asp Phe Phe Val
115      120      125
Met Ser Asp Tyr Arg Gly Phe Gly Ile Gly Ser Glu Ile Leu Lys Asn
130      135      140
Leu Ser Gln Xaa Val Ala Met Arg Cys Arg Cys Ser Ser Met His Phe
145      150      155      160
Leu Val Ala Glu Trp Xaa Xaa Xaa Xaa Asn Glu Pro Ser Ile Asn Phe
165      170      175
Tyr Lys Arg Arg Gly Ala Ser Asp Leu Ser Ser Glu Glu Gly Trp Xaa
180      185      190

```

SUBSTITUTE SHEET (RULE 26)

100

Xaa Xaa Xaa Xaa Arg Leu Phe Lys Ile Asp Lys Glu Tyr Leu Leu Lys
 195 200 205
 Met Ala Thr Glu Glu Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 210 215 220
 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:78:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:78:

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Met
 1 5 10 15
 Asn His Ala Gln Leu Arg Arg Val Thr Ala Glu Ser Phe Ala His Tyr
 20 25 30
 Arg His Gly Leu Ala Gln Leu Leu Phe Glu Thr Val His Gly Gly Xaa
 35 40 45
 Xaa Ala Ser Val Gly Phe Met Ala Asp Leu Asp Met Gln Gln Ala Tyr
 50 55 60
 Ala Trp Cys Asp Gly Leu Lys Ala Asp Ile Ala Ala Gly Ser Leu Leu
 65 70 75 80
 Leu Trp Val Val Ala Xaa Xaa Xaa Xaa Glu Asp Asp Asn Val Leu
 85 90 95
 Ala Ser Ala Xaa Xaa Gln Leu Ser Leu Cys Gln Lys Pro Asn Gly Leu
 100 105 110
 Asn Arg Xaa Xaa Xaa Xaa Xaa Xaa Ala Glu Val Gln Lys Leu Met Val
 115 120 125
 Leu Pro Ser Ala Arg Gly Arg Gly Leu Gly Arg Gln Leu Met Asp Glu
 130 135 140
 Val Glu Gln Xaa Val Ala Val Lys His Lys Arg Gly Leu Leu His Leu
 145 150 155 160
 Asp Thr Glu Ala Xaa Xaa Xaa Xaa Gly Ser Val Ala Glu Ala Phe
 165 170 175
 Tyr Ser Ala Leu Ala Tyr Thr Arg Val Gly Glu Leu Pro Gly Tyr Cys
 180 185 190
 Ala Thr Pro Asp Gly Arg Leu His Pro Thr Ala Ile Tyr Phe Lys Thr
 195 200 205

SUBSTITUTE SHEET (RULE 26)

101

Leu Gly Gln Pro Thr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 210 215 220
 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:79:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 240 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:79:

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 1 5 10 15
 Xaa Xaa Xaa Xaa Met Pro Asn Val Thr Ile Ala Arg Glu Ser Pro Leu
 20 25 30
 Gln Asp Ala Val Val Gln Leu Ile Glu Glu Leu Asp Arg Xaa Xaa Xaa
 35 40 45
 Xaa Xaa Xaa Xaa Xaa Tyr Leu Gly Asp Leu Tyr Pro Ala Glu Ser Asn
 50 55 60
 His Leu Xaa Xaa Xaa Leu Asp Leu Gln Thr Leu Ala Lys Pro Asp Ile
 65 70 75 80
 Arg Phe Leu Val Ala Xaa Xaa Xaa Xaa Xaa Arg Arg Ser Gly Thr Val
 85 90 95
 Val Gly Cys Xaa Xaa Gly Ala Ile Ala Ile Asp Thr Glu Gly Gly Tyr
 100 105 110
 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Gly Glu Val Lys Arg Met Phe Val
 115 120 125
 Gln Pro Thr Ala Arg Gly Gly Gln Ile Gly Arg Arg Leu Leu Glu Arg
 130 135 140
 Ile Glu Asp Xaa Glu Ala Arg Ala Ala Gly Leu Ser Ala Leu Leu Leu
 145 150 155 160
 Glu Thr Gly Val Tyr Xaa Xaa Xaa Xaa Gln Ala Thr Arg Ile Ala Leu
 165 170 175
 Tyr Arg Lys Gln Gly Phe Ala Asp Arg Gly Pro Phe Gly Pro Tyr Gly
 180 185 190
 Pro Asp Pro Leu Ser Leu Phe Met Glu Lys Pro Leu Xaa Xaa Xaa Xaa
 195 200 205
 Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 210 215 220

SUBSTITUTE SHEET (RULE 26)

102

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 225 230 235 240

(2) INFORMATION FOR SEQ ID NO:80:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:80:

Xaa Xaa Xaa Xaa Xaa Met Pro Ile Asn Ile Arg Arg Ala Thr Xaa Ile
 1 5 10 15
 Asn Asp Ile Ile Cys Met Gln Asn Ala Asn Leu His Asn Leu Pro Glu
 20 25 30
 Asn Tyr Met Met Lys Tyr Tyr Met Tyr His Thr Leu Ser Trp Pro Glu
 35 40 45
 Ala Ser Phe Val Ala Thr Thr Thr Thr Leu Asp Cys Glu Asp Ser Asp
 50 55 60
 Glu Gln Asp Glu Asn Asp Lys Leu Glu Leu Thr Leu Asp Gly Thr Asn
 65 70 75 80
 Asp Gly Arg Thr Ile Lys Leu Asp Pro Thr Tyr Leu Ala Pro Gly Glu
 85 90 95
 Lys Leu Val Xaa Xaa Gly Tyr Val Leu Val Lys Met Asn Asp Asp Pro
 100 105 110
 Asp Gln Gln Asn Glu Pro Pro Asn Gly His Ile Thr Ser Leu Ser Val
 115 120 125
 Met Arg Thr Tyr Arg Arg Met Gly Ile Ala Glu Asn Leu Met Arg Gln
 130 135 140
 Ala Leu Phe Ala Leu Arg Glu Val His Gln Ala Glu Tyr Val Ser Leu
 145 150 155 160
 His Val Arg Gln Ser Xaa Xaa Xaa Xaa Asn Arg Ala Ala Leu His Leu
 165 170 175
 Tyr Arg Asp Thr Leu Ala Phe Glu Val Leu Ser Xaa Xaa Xaa Xaa Ile
 180 185 190
 Glu Lys Ser Tyr Tyr Gln Asp Gly Glu Asp Ala Tyr Ala Met Lys Lys
 195 200 205
 Val Leu Lys Leu Glu Glu Leu Gln Ile Ser Asn Xaa Xaa Xaa Phe Thr
 210 215 220
 His Arg Arg Leu Lys Glu Asn Glu Glu Lys Leu Glu Asp Asp Leu Glu

SUBSTITUTE SHEET (RULE 26)

103

225

230

235

240

(2) INFORMATION FOR SEQ ID NO:81:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 240 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:81:

```

Met Glu Ile Val Tyr Lys Pro Leu Asp Ile Arg Asn Glu Glu Gln Phe
 1           5           10           15

Ala Ser Ile Lys Lys Leu Ile Asp Ala Asp Leu Ser Glu Pro Tyr Ser
 20           25           30

Ile Tyr Val Tyr Arg Tyr Phe Leu Asn Gln Xaa Xaa Xaa Trp Pro Glu
 35           40           45

Leu Thr Tyr Ile Ala Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 50           55           60

Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Val Asp Asn Lys Ser
 65           70           75           80

Gly Thr Pro Asn Ile Pro Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
 85           90           95

Xaa Xaa Ile Xaa Xaa Gly Cys Ile Val Cys Lys Met Asp Xaa Xaa Xaa
100           105           110

Pro His Arg Asn Val Arg Leu Arg Gly Tyr Ile Gly Met Leu Ala Val
115           120           125

Glu Ser Thr Tyr Arg Gly His Gly Ile Ala Lys Lys Leu Val Glu Ile
130           135           140

Ala Ile Asp Lys Met Gln Arg Glu His Cys Asp Glu Xaa Ile Met Leu
145           150           155           160

Glu Thr Glu Val Glu Xaa Xaa Xaa Xaa Asn Ser Ala Ala Leu Asn Leu
165           170           175

Tyr Xaa Glu Gly Met Gly Phe Ile Arg Met Lys Xaa Xaa Xaa Xaa Arg
180           185           190

Met Phe Arg Tyr Tyr Leu Asn Glu Gly Asp Ala Phe Lys Leu Xaa Xaa
195           200           205

Ile Leu Pro Leu Thr Glu Lys Ser Cys Thr Arg Ser Thr Phe Leu Met
210           215           220

His Gly Arg Leu Ala Thr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
225           230           235           240

```


104

(2) INFORMATION FOR SEQ ID NO:82:

- (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 240 amino acids
 (B) TYPE: amino acid
 (C) STRANDEDNESS: single
 (D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:82:

Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	1	5	10	15
Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	20	25	30	
Met	Asn	Tyr	Gln	Ile	Val	Asn	Ile	Ala	Glu	Cys	Ser	Asn	Tyr	Gln	Leu	35	40	45	
Glu	Ala	Ala	Asn	Ile	Leu	Thr	Glu	Ala	Phe	Asn	Asp	Leu	Gly	Asn	Asn	50	55	60	
Ser	Trp	Pro	Asp	Met	Thr	Ser	Ala	Thr	Lys	Glu	Val	Lys	Glu	Cys	Ile	65	70	75	80
Glu	Ser	Pro	Asn	Leu	Cys	Phe	Gly	Leu	Leu	Ile	Asn	Asn	Ser	Leu	Val	85	90	95	
Gly	Trp	Ile	Xaa	Xaa	Gly	Leu	Arg	Pro	Met	Tyr	Lys	Glu	Thr	Trp	Glu	100	105	110	
Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Leu	His	Pro	Leu	Val	Val	115	120	125	
Arg	Pro	Asp	Tyr	Gln	Asn	Lys	Gly	Ile	Gly	Lys	Ile	Leu	Leu	Lys	Glu	130	135	140	
Leu	Glu	Asn	Arg	Xaa	Ala	Arg	Glu	Gln	Gly	Ile	Ile	Gly	Ile	Ala	Leu	145	150	155	160
Gly	Thr	Asp	Asp	Glu	Tyr	Tyr	Arg	Thr	Ser	Leu	Ser	Leu	Ile	Thr	Ile	165	170	175	
Thr	Glu	Asp	Asn	Ile	Phe	Asp	Ser	Ile	Lys	Asn	Ile	Lys	Asn	Ile	Asn	180	185	190	
Lys	His	Pro	Tyr	Glu	Phe	Tyr	Gln	Lys	Asn	Gly	Tyr	Tyr	Ile	Val	Gly	195	200	205	
Ile	Ile	Pro	Asn	Ala	Asn	Gly	Lys	Asn	Lys	Pro	Asp	Ile	Trp	Met	Trp	210	215	220	
Lys	Ser	Leu	Ile	Lys	Glu	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	225	230	235	240

SUBSTITUTE SHEET (RULE 26)

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WHAT IS CLAIMED IS:

1. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequences selected from the group consisting of SEQUENCE ID NOS: 1 and 2.

2. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 3.

3. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequences set forth in SEQUENCE ID NO: 4.

4. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequence set forth in SEQUENCE ID NO: 5.

5. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 6.

6. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequence set forth in SEQUENCE ID NO: 7.

7. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 8.

8. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequence set forth in SEQUENCE ID NO: 9.

9. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 10.

10. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequence set forth in SEQUENCE ID NO: 11.

11. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 12.

12. An isolated nucleic acid sequence comprising the nucleic acid sequence encoding the sequences selected from the group consisting of SEQUENCE ID NO: 14 and 15.

13. An isolated protein sequence comprising the amino acid sequence set forth in SEQUENCE ID NO: 16.

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14. A DNA sequence comprising a sequence complementary to an isolated nucleic acid sequence of claim 1.

15. A transformed plant cell comprising the nucleic acid sequence selected from the group consisting of SEQUENCE ID NOS: 1, 2, 4, 5, 7, 9, 11, 14, and 15.

16. A plant comprising a heterologous nucleic acid sequence selected from the group consisting of SEQ ID NOS: 1, 2, 4, 5, 7, 9, 11, 14, and 15.

17. A DNA sequence comprising a sequence complementary to an isolated nucleic acid sequence selected from the group consisting of SEQ ID NOS: 1, 2, 4, 5, 7, 9, 11, 14, and 15.

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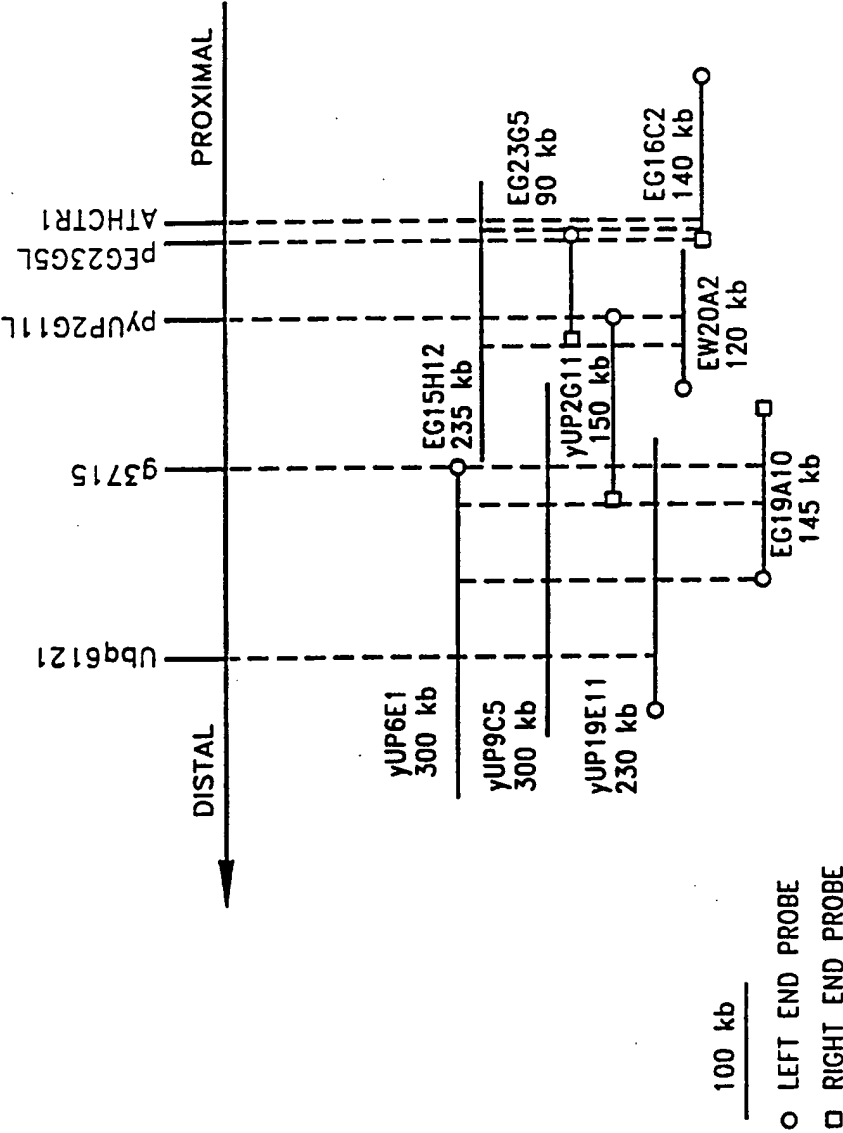


FIG. 1

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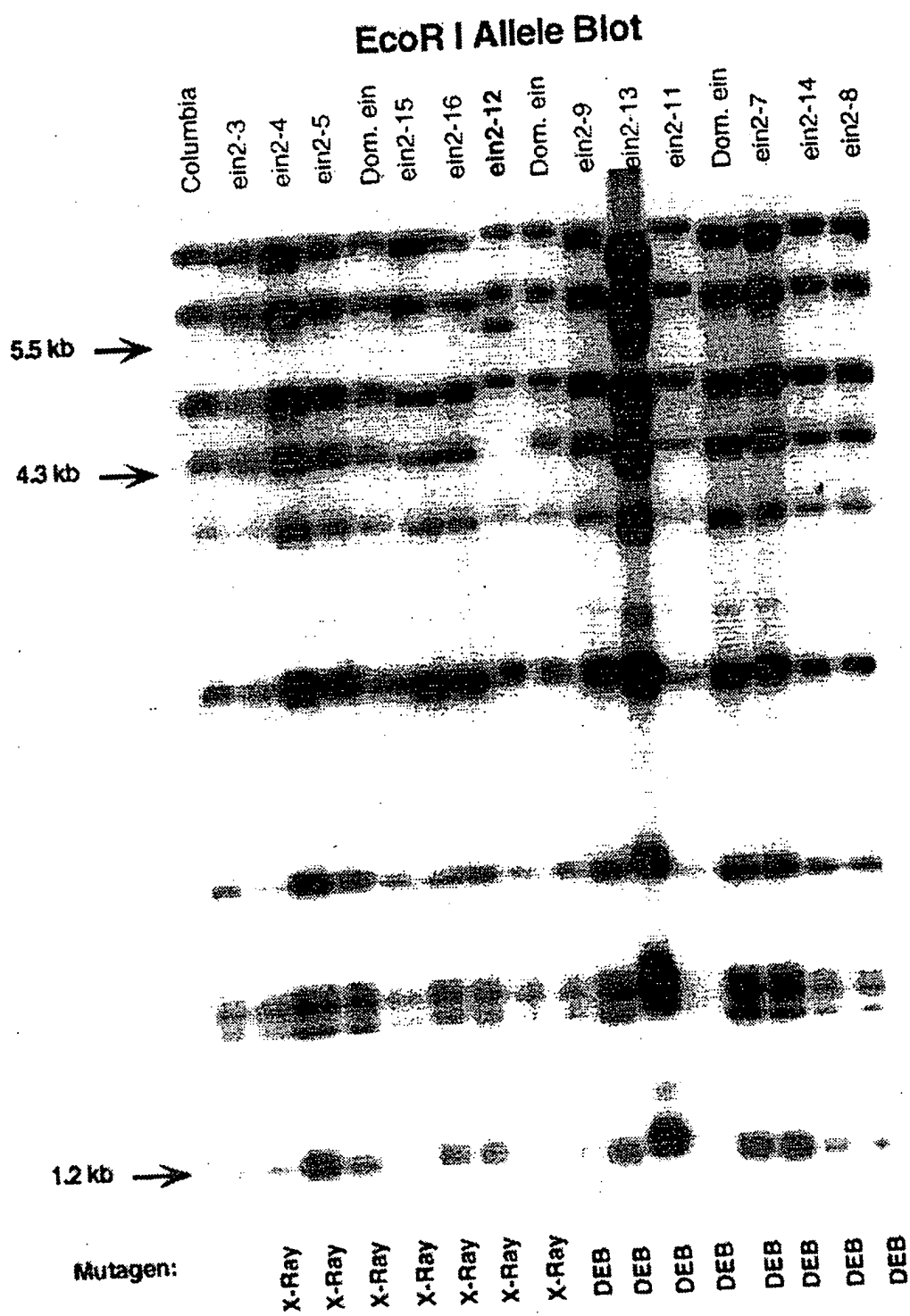
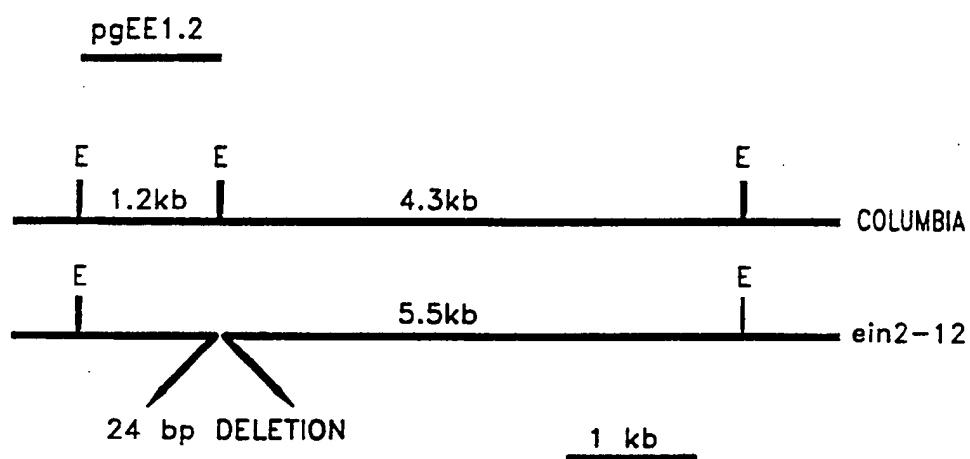


FIG. 2

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*FIG. 3*

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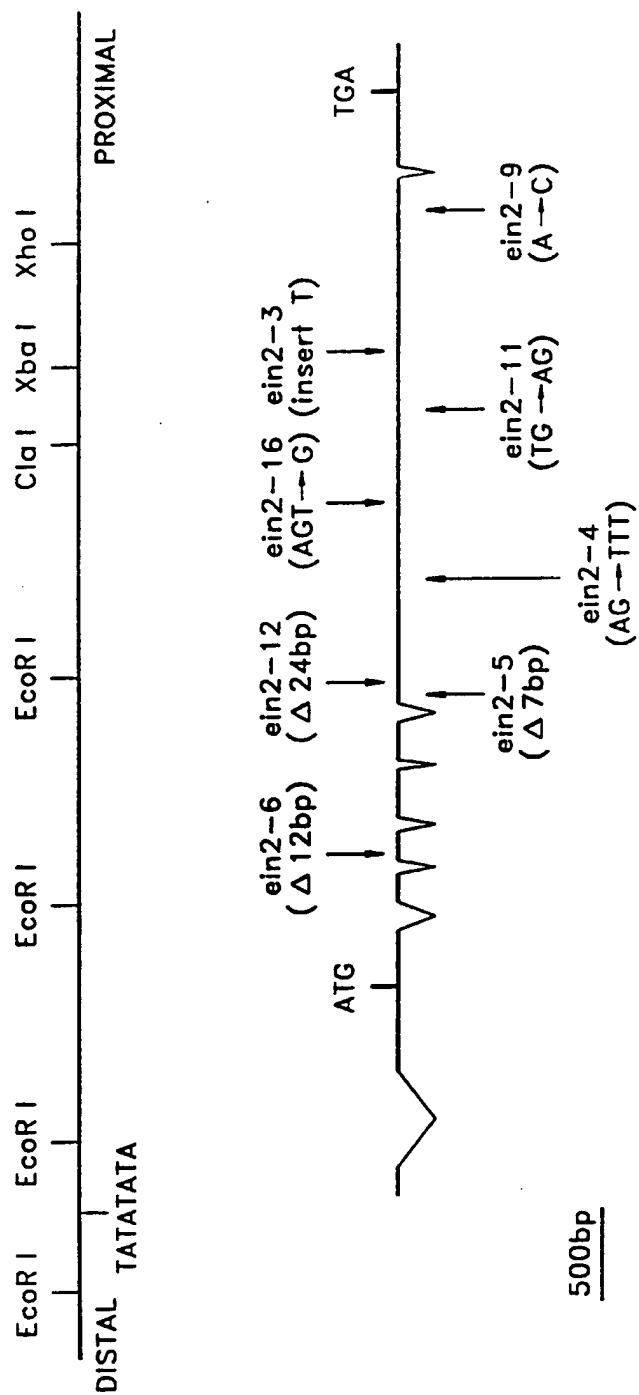


FIG. 4

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[illegible]

FIGURE 5a

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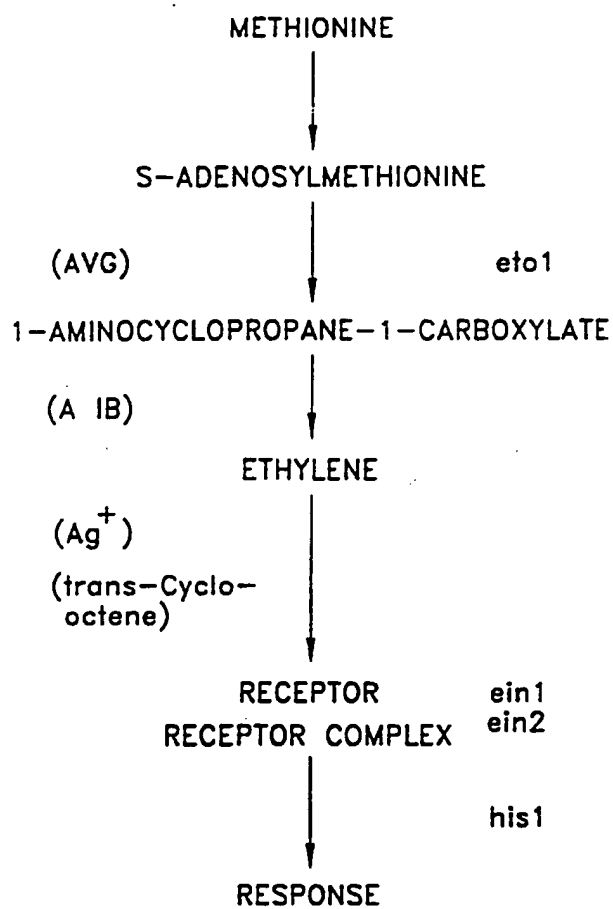
CCGATCAATATAGACAACACGCTTCTTCTAATGTTGATGCGAGGCTAAGCTTCTTACGCTTCAGGCACCTGTATTTCTAAGACTTATTAACCTTGAAGGATCCGA
PTGINTTGTGACAAACGATGGAGTTGATGAAGAACTGATTCACCGGTAGCTGCACGAGAGAGTTTATCTATGAGCTGAAGCTTCGANAATNAAACCAAGTGG
W L F G Q S D G V D E E L I D R V A A R E K F I Y E A E A E I N Q V G
GTACATATGGGGGACCACTAATTCATCGGTCTCTAAGTGGAGATGGTTCGTTGGACAGCTGATTTGATGTAGCTTTTGGACTTTGGACATTTGGTGCACTTCCACCGTGT
H M G E P L I S S V P N C G D G C V W R A D L I V S F G V W C I H R V

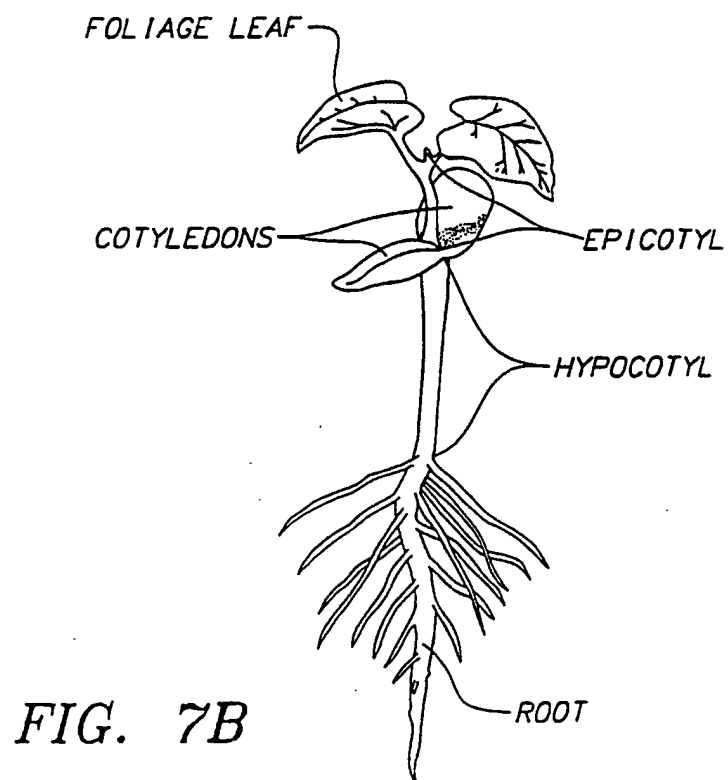
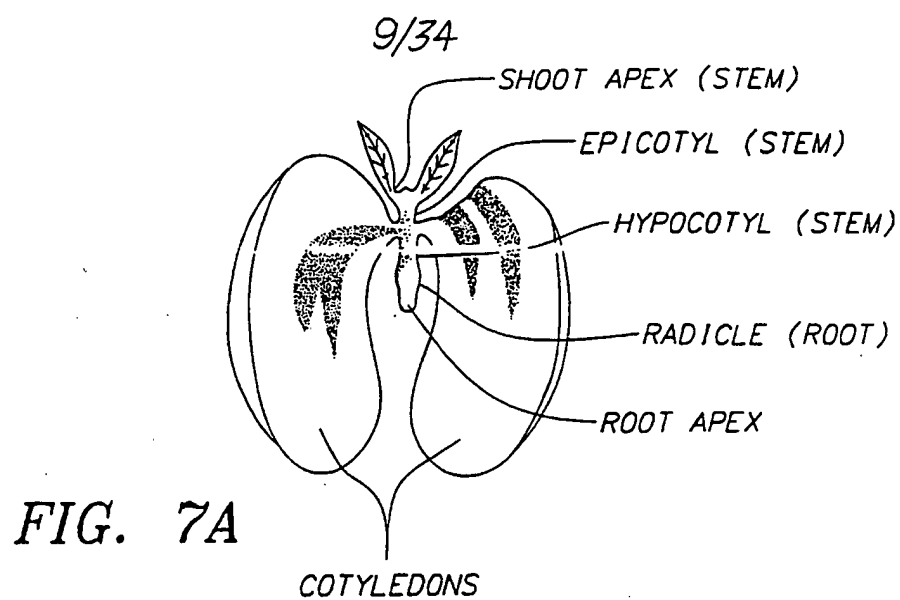
CTTGACTTCTCTCATGAGAGCGGCTTTTGGGGAAGTACACTACGTTCTCAACCGCTACAGGTAcaaaaaacccgacgtagttcataggaaatcac
L D L S L M E S R P E L W G K Y T V Y L N R L Q G
agttttgcagtttgaaatatattacatgtatgatttataaacagcagtgattgattgattcgcgggttctcaaaagctgcggacacccaatgacacacgctgcttttgcctttcagat
v i d p a f s k l r t p t m t p c f c l o i
TCAGCGCAGCCACCGAGAGCGAGTCCGACTTCAGCTAACGGAATGTACCTCCGCTGCAAAACCGCTAAGGCCAAATGCACACCGCTGCACACTTCTTGATC
P A S H Q R A S P T S A N G M L P P A A K P A K G K C T A V T L L D L
TAATCAAAAGCTTGAAATGGCAATCTTGTAGAAAGCCGGAACCGGTACAGCTGCAGGTATGTGCTTCCCAAAGGGGAAGAGAATTTGGCTTCGGTTTCG
I K D V E M A I S C R K R T G T A G D V A F P K G K E N L A S V S
AAAGCGGTATAAACGTCCGTTATCGAATAAACCACTAAGGTATGAATCAGGATGGACCCGGTTCAAGAAAACCGTACGTGCTGAGATCATTTGGTGTAGSAAAGAA
K R Y K R R L S N K P Y E S G W T F C K K R D C V R I G L K K K
GAACTATGTGACAAATCTCAAGATGCAATCGACGTGAGAGGGAACCGAAGAACTCAAAACTCTGCTTCTGATTTGATTCCTCTCTCTCTGTTTGTATTGATTAAAGAA
N I V R N L M I K V T S R G K P K N Q N S R F +
PAGAAGAAAATAATGGATTTTTCGCTCTTTTTCTTCTCATGAAATTTTGGTGTGTAATGTATTGTTTATATACATATATCATCATATAGGACCAT
AGCTACAAACCGAATCTTGTGTAATTTCTATGCGAATACGAAGAATTCGTG

Figures 5a, 5b and 5c: The sequence of the EIN2 locus.

FIGURE 5c

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*FIG. 6*



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	1		50
pileup.msf(ei11)	..hhhmMMFM	EMGMYGND FSSs..Tald	vCPIQaEqE pVVeDvDYtD
pileup.msf(ei13)	iiitttMMFN	EMGMCGND FSSgSLgEVD	fCPvPQaEpD alVED.DYtD
pileup.msf(ei12)	..dsmdMynN	niGMFrSLvc sSappFTegh	MCs...dsht alcDDIs.sD
pileup.msf(ei13)mg	DLaM.....SvaDir MenePddlas dnVaEIDvaD
Consensus	-----M-----	-----S-----	-----D-----
	51		100
pileup.msf(ei11)	DEmDVDELEk	RMWRDKMRLK RLKEQqSkKc	EGVDgsKQRO SW..EQARRK
pileup.msf(ei13)	DEiDVDELEr	RMWRDKMRLK RLKEQd.KGK	EGVDQoKQRO SQ..EQARRK
pileup.msf(ei12)	EEeEIEELEk	kiWRDKqRLK RLKEmoKnGl	gtrlllKQqh ddfpEhsskr
pileup.msf(ei13)	EEiDoDDLEr	RMWkDrvRLK RiKErOKaGs	qGaqt..Ketp kkiSDQAqRK
Consensus	-E-----LE-	--W-D--RLK R-KE-----	-----K----
	101		150
pileup.msf(ei11)	KMSRAQDGIL	KYMLKMEVC KAQGFVYGII	PEkGKPVtGa SDNLRWWKD
pileup.msf(ei13)	KMSRAQDGIL	KYMLKMEVC KAQGFVYGII	PEngKPVtGa SDNLRWWKD
pileup.msf(ei12)	tMykaQDGIL	KYMsKtMErY KAQGRVYGIV	lEnGktVaGs SDNLRWWKD
pileup.msf(ei13)	KMSRAQDGIL	KYMLKLMEVC KvrGFVYGII	PEkGKPVaGs SDNiRoWwKE
Consensus	-M--AQDGIL	KYM-K-ME-- K--GFVYGII-	-E-GK-V-G- SDN-R-WWk-
	151		200
pileup.msf(ei11)	KVRFDNRNGPA	AlAKYQsENN lSGGSnDcNs	lVGPTPHTLQ ELQDTTLGSL
pileup.msf(ei13)	KVRFDNRNGPA	AltKYQaENN lp.GiHEGNN	plGPTPHTLQ ELQDTTLGSL
pileup.msf(ei12)	KVRFDNRNGPA	AliKhQrDiN lSdGSDsGse	vgdsIaqlLI ELQDTTLGaL
pileup.msf(ei13)	KVvFDkNGPA	AlAKYeeEcl afGkSDgnrNsqfvLQ DLQDaTLGSL
Consensus	KV-FD-NGPA	Al-K-----	-----L- -LQD-TLG-L
	201		250
pileup.msf(ei11)	LSALMQHCDP	PQRRFPLEKG VsPPWWPnGn	EEWWPQLGLP nE..QGPPPY
pileup.msf(ei13)	LSALMQHCDP	PQRRFPLEKG VPPWWPnGk	EDWWPQLGLP KD..QGPaPY
pileup.msf(ei12)	LSALfPhCnP	PQRRFPLEKG VtPPWWPtGk	EDWWdQLsLP vDfrgvPPPY
pileup.msf(ei13)	LSsLMQHCDP	PQRkYPLEKG tPPPWWPtGn	EEWWvkLGLP Ks...qsPPY
Consensus	LS-L--HC-P	PQR--PLEKG --PPWWp-G-	E-WW--L-LP -----PY
	251		300
pileup.msf(ei11)	KKPHDLKKoW	KVGVLTAIVK HMsPDIAKIR	KLVRQSKCLQ DKMTAKESAT
pileup.msf(ei13)	KKPHDLKKoW	KVGVLTAIVK HMFpDIAKIR	KLVRQSKCLQ DKMTAKESAT
pileup.msf(ei12)	KKPHDLKKIW	KIGVLigVlr HMasDlsnlp	nLVRrSrSLQ EKMTsrEgAl
pileup.msf(ei13)	rKPHDLKKmW	KVGVLTAIVn HMLPDIAKIk	rhVRWSKCLQ DKMTAKESAi
Consensus	-KPHDLKK-W	K-GVL--Vl- HM--DI--l-	--VR-S--LQ -KMT--E-A-
	301		350
pileup.msf(ei11)	WLAliNQEEv	vaReLYPES.CPPLSs	SsslGSgSLL iNDCEsYDVE
pileup.msf(ei13)	WLAliNQEEs	laReLYPES.CPPLSL	Sg..GSsSLL mNDCSqYDVE
pileup.msf(ei12)	WLAalyrEka	ivdq.....ioM SrenntSnF lvpotggDpD
pileup.msf(ei13)	WLAVINQEES	liqqssDng nsnvtethrr	gnnadrrkpV vNsdsDYDvD
Consensus	WLA---E--	-----	-----D--

FIG. 8A

FIG. 8

FIG. 8A
FIG. 8B

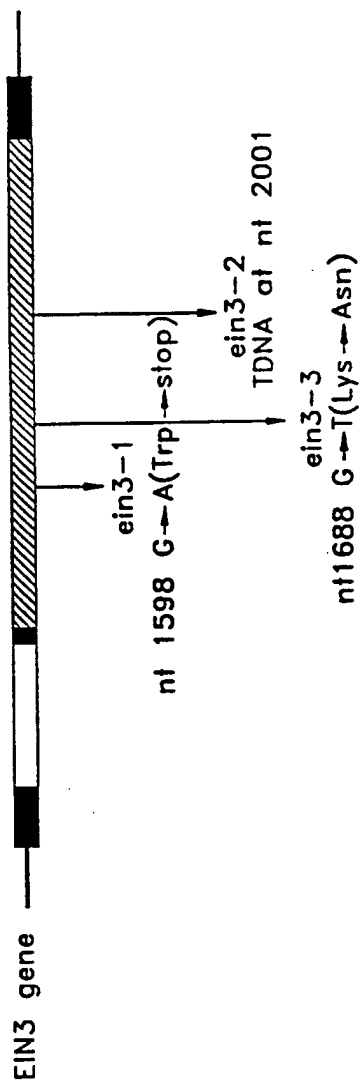
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	351		400
pileup.msf(ei11)	GFEKEqHgFD VEErKPEiVM mhpLasfgVA KMQhFPIKEE VattvNIEFT		
pileup.msf(ei13)	GFEKESH.YE VEEIKPEkVM nssnfGm.VA KMhdFPVKEE Vpag.NsEFm		
pileup.msf(ei12)	vLfpEstdYD VE..... ..LiGgthr tnQqYP...E fennyNcvYk		
pileup.msf(ei13)	GtEeaSgsvs skDsrrnql.q KeQptalsvs VrdqdkakEk		
Consensus	-----	-----	-----
	401		450
pileup.msf(ei11)	RKRKqNnDMN vmVMDSagY TCENgqCPHS kmnLGFqDRs SRDNHQMvCP		
pileup.msf(ei13)	RKRKpNRDLN t.IMDR.TvF TCENlgCaHS eISRGFDRN SRDNHQLaCP		
pileup.msf(ei12)	RKfeedfgMp m....hpTIL TCENslCPyS QphMGFLDRN IRENHQMlCP		
pileup.msf(ei13)	RrRKrpR...iRSgtv nrqeeeqPea QqrniLpDmN hvDopILeYn		
Consensus	R-----	-----	-----D-----
	451		500
pileup.msf(ei11)	YRDnRLaYGA ..SkFHMgGm KIVV...pqq PV.....QPI DLsGVgVPEn		
pileup.msf(ei13)	hRDsRLpYGA apSrFHvnev KpVVgFpqPr PVnsva.QPI DLTGI.VPED		
pileup.msf(ei12)	YkvTsF....yqpT.kPy gMTGIMVP..		
pileup.msf(ei13)	ingThqeddv vdpniaLGe dngeLvvPe fnNnyTyIPi vneqtMmPvD		
Consensus	-----	-----	-----P-----P-----
	501		550
pileup.msf(ei11)	GQKMItELma MYDRnVQS.. ..nQTpptLM ENQSmvidak aaqNqQInFn		
pileup.msf(ei13)	GQKMIsELms MYDRnVQS.. ..nQT.amvM ENQSVslLqP tvhNhQehLq		
pileup.msf(ei12)cpDyng M.qqqVQS.. ..fQdqf... .NhpnDlyrP kapqr.....		
pileup.msf(ei13)	erpMlygpnq nqElqfgSgy nfynpsavFv hNQedDiLht qie.....		
Consensus	-----	-----S-----	-----N-----
	551		600
pileup.msf(ei11)SGNQm Fmq..... ..		
pileup.msf(ei13)	fpgrnmvegsf fednipnra NnnnsSnNQt Ffqgnnnnnn vFkFdaDhn		
pileup.msf(ei12)GNdd Lved..... ..		
pileup.msf(ei13)m NtqapphNag FeeapggvIq pLgLIgnEdg		
Consensus	-----	-----N-----	-----
	601		650
pileup.msf(ei11)qgtN nGVNNRFQMV FDSTpFDMAa FDYRDDWqtG amEgmGkqqq		
pileup.msf(ei13)	nfeoaHnnN nssgNRFQLV FDSTpFDMAa FDYRDDmSmp Gv..VGTmdg		
pileup.msf(ei12)LNpsp stINqnLgLV L.pTdFn... ..G GeEtVGTenn		
pileup.msf(ei13)	vtgseLpqyq sGllspL... ..TdLDfdy ggFgDDFSwf Ga.....		
Consensus	-----	-----T-----	-----
	651	664	
pileup.msf(ei11)	qQQQQQDVSI W...		
pileup.msf(ei13)	MQQkQQDVSI W...		
pileup.msf(ei12)	LhnQgQEIpI swiq		
pileup.msf(ei13)		
Consensus	-----	-----	

FIG. 8B

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 CODING REGION
 INTRON
 NON-TRANSLATED REGIONS



PREDICTED POLYPEPTIDE
 628 aa


 acidic basic Asn repeats

FIG. 9

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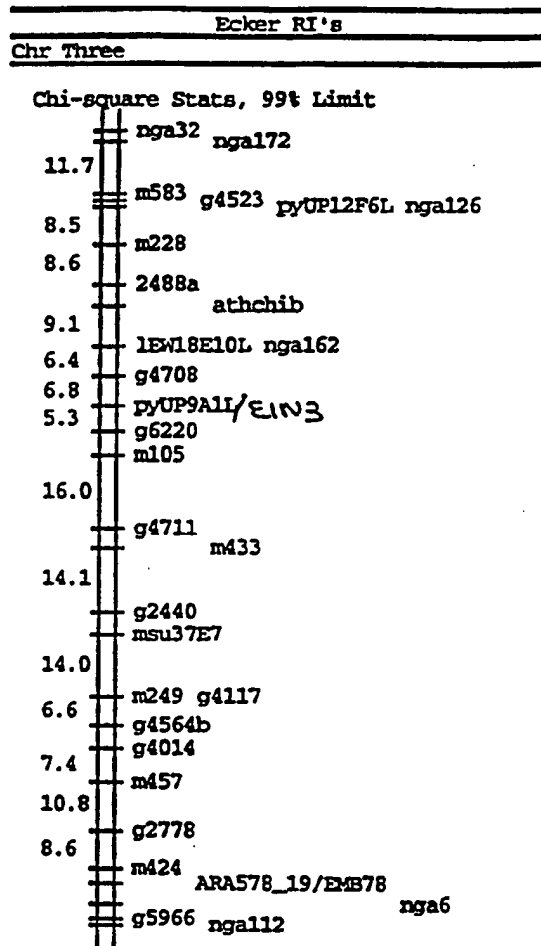


FIGURE 10

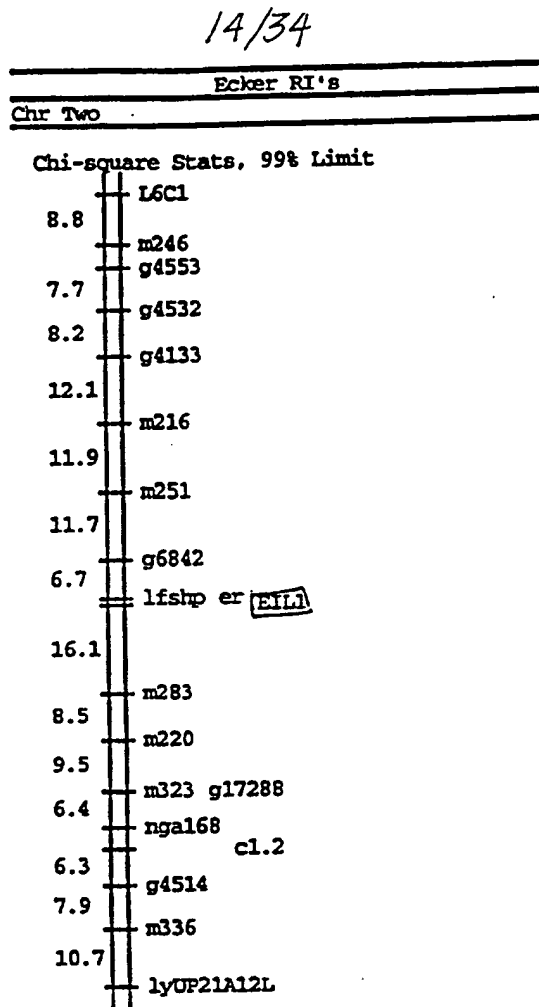


FIGURE 11

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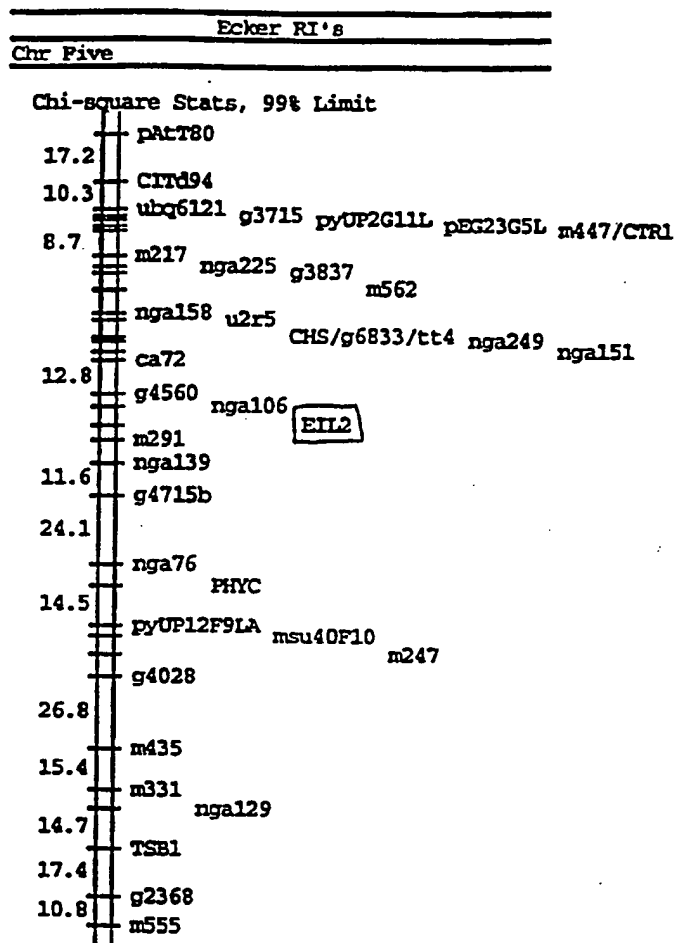


FIGURE 12

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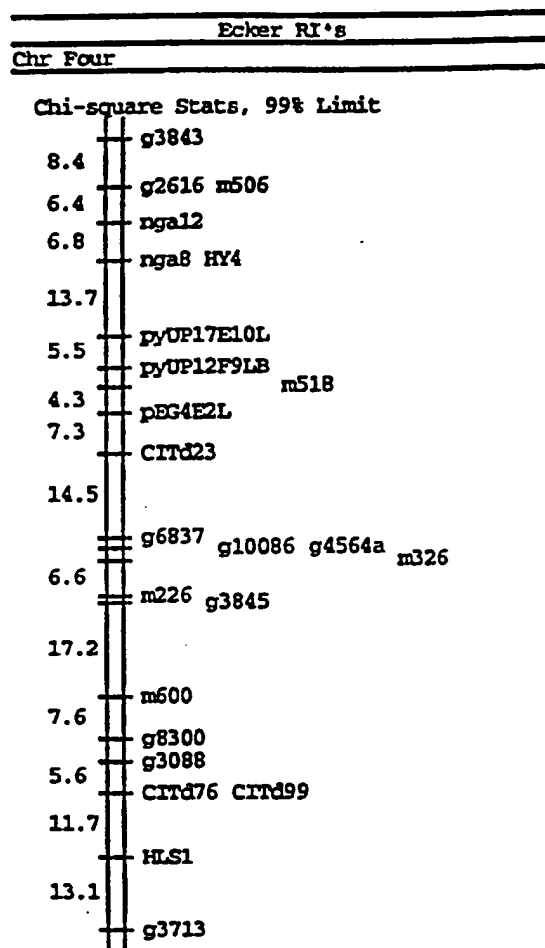


FIGURE 13

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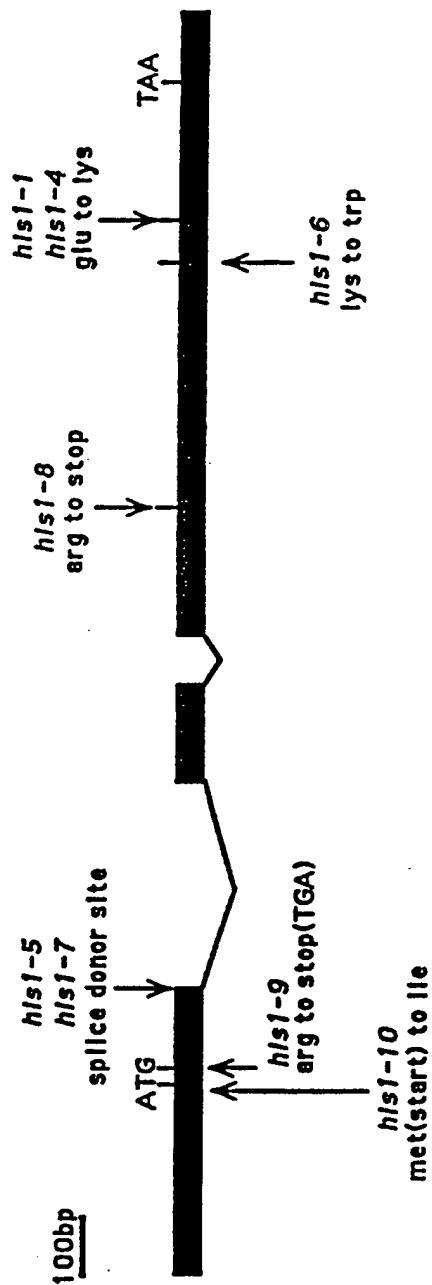


FIGURE 14

1mfgyrsnvkvrrlttdrlvrlv hDr-daWrlad YyoenrhFlk pweprDEsh cypsgwqarImletik vseslelhav aEnhvkvplyq LicknktWlq qslnwpqfvq seedtrktlvqmlw ssndvtqags rpktklggs. msilatVktg pDeisamrav LdIFgkEFed iplysdrqpt neylonlIhsmlr ssndvtqags rpktklggs mgjirtctlg pDqvksmraa LdIFgrEFgd valysqhqp sdylgnlIrs mttlddtoyr yrtsvpgdae aiealdgsft tdtvfrvtat gDgflrevp vdppltkv... fpddesDDes ddgedgdps mttthgstye frsarpgdae aiealdgsft tsvfevdt gDgflrevp adpplkv... fpddgsDge dgaegedadsmk isvipeqvae tlda.enhfi vrevfVhLS dqgfelstrs vspYrkDY... isdddsDE... ..dsmakfk irpataadcsdiIrLi kElakYeyme dqviltEkdl qedgfgEhpf yhcIvaevpkmakfv irpataadcsdiIrLi kElakYeyme eqviltEkdl ledgfgEhpf yhcIvaevpkmhaq lrrvtasfa hyrhglaql fEtvhgg..a svqfmaDLdm qqayawcDgl kadiagssIImpnvtiares plqadvqLi eEldr..... YlgDlyp aeshl...I dlqtlakpdimpini rrot.indii cmqnanlhnI penymkyYm yhtIsWpeas FvattttLdc edsdeqDEnd kleltldgtN meivykpldi rneeqfasik klidadlsep ysiyvyryfl nq...Wpelt Yia.....vdnks mtvvreydpt rdlvgvedve rncevgsqk lsiftdlldg picriRhspS YlmLvaEmgt e...kkEivg mirgciktvtmk..... ..mnyqlvni aEcsnYqlea onILteafnd lgnswpDmt satkevkeci -----v-L- -E-----F--EF-- --D--	80
160	gminefhkqg safyfglFdp dekeiligvan fsnvrgsfh aCylgYslgq kwqGkGlmfe altaairyma rtqhihrimo gnv.mlharg yakmfmiF.. kedeligvis f.nriepInk taiegYwlde shqGqGllsq alqaLihyA qsgelrrfvi etFIAlaafd rgtaiaggLA. .aYVlpkfeq arse..... iYlvdLaVas shRRlGVata LishLkr.vA velGayviyv ktFIAlaafd qeavvgalA. .aYVlpkfeq arse..... iYlvdLaVas eHRRqGlatO LinLKh.eA nalGayviyv rtFvAygD..dgdLA. .GFVvisysa wnrr..... ltVedieVap eHRGhGVGrO lMglate.fA gerGaghIwL rtFvAvgA..dgdLA. .GFaavsysa wnqr..... ltledieVap gHRGkGIGrv lMrhaad.fA rerGaghIwL acYgAf.i...dqeLv. .GkleIn.st wndl..... astehivVsh lHRGkGVahs Liefakk.wA lsrqllgIrL ehWtp..... eghsivgFA. .mYyFLydpw igkl..... lYledFfVms dyRGfGIGse ilknLsq.vA mkcrccssmhF	81
{rimJ: E.coli} {rimI: E.coli} {N3nat: Pseudomonas} {Nnat: E.coli} {natI: Streptomyces} {sat: Streptomyces} {sat: E.coli} {ssat: Mouse} {ssat: Human} {tab: Pseudomonas} {tab: Azospirillum} {ardI: Yeast} {MAK3: Yeast} {HLSI: Arabidopsis} {oac(6?):Citrobacter} Consensus	{rimJ: E.coli} {rimI: E.coli} {N3nat: Pseudomonas} {Nnat: E.coli} {natI: Streptomyces} {sat: Streptomyces} {sat: E.coli} {ssat: Mouse}	

FIG. 15

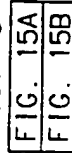


FIG. 15A

240

161

```
{rimJ: E.coli}
{rimI: E.coli}
{N3nat: Pseudomonas}
{Nnat: E.coli}
{natI: Streptomyces}
{sat: Streptomyces}
{ssat: E.coli}
{ssat: Mouse}
{ssat: Human}
{tab: Pseudomonas}
{tat: Azospirillum}
{ardI: Yeast}
{MAK3: Yeast}
{HLSI: Arabidopsis}
{aac(6'): Citrobacter}
Consensus
```

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N krsqdLlArI GFekeGyakd yllidgWrd hvltalTtpd wtpgr
 N pqsqvalrn GFileGclq oeflndotdd vnlYariids q
 d dPAValYtkl Gvredvmhfd idprtat
 d dPAValYtkl Gireevmhfd idpstat
 N aPAIhaYrrm GfTlcGldta lydgatasdge rqlYMsmcp p
 N aPAIhaYrrm GFafcGldsa lyqgtasege .halYMsmcp p
 N vPACnLYakc GfTlgGldf tyktrpqvsn etamYwywfs gaqdda
 N ePsnfYkrr Gasdlsseeg w...rLfk idkeYLlkma oee
 N ePsnfYkrr Gasdlsseeg w...rLfk idkeYLlkma tee
 g svAeafYsal GYtrvGelpg ycalpdgrlh ptaiYfklng qpt
 q atrlndYrk GfadrGpfpg ygdpdlsfsm ekpl
 N raAlhLYrdt lafevls... .leksyyqdg edaYalmkkvl kleeqlqsn. .flhrr lke neekleddle
 N saAlnLYeg mgfirmlk... .rmfryylne gdafKL.il pleteksctrs tflmhgr lot
 N qasVnLftg cgysfrltps ilvnpvyahr vnvsvrrvtvi klepvdaet. .lyriifsl teff
 gtddeyyrls lslititedn inkhpvryq knGYivvgii pnongknkpd imwmks lke
 N -PAI-LY--- GF---G---Y

FIG. 15B

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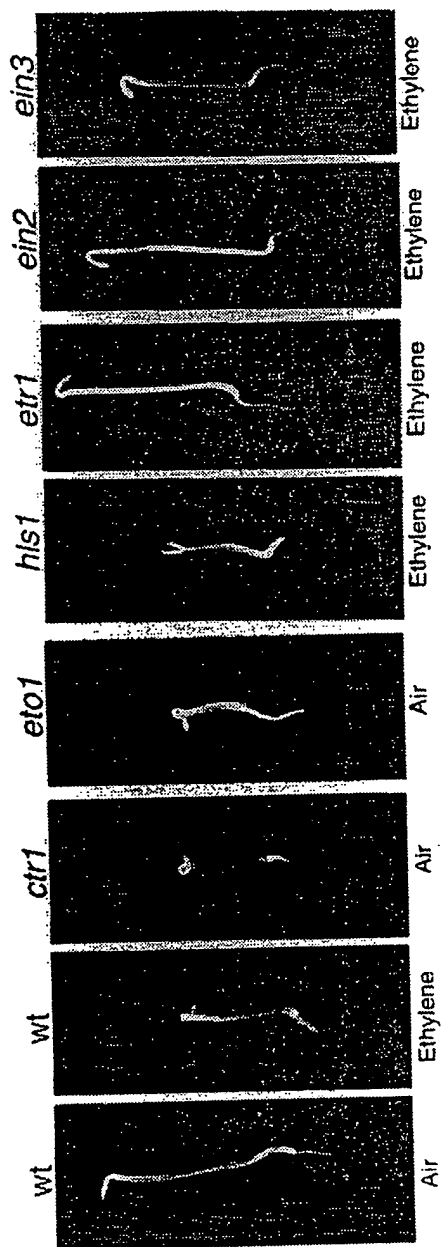


FIG. 16

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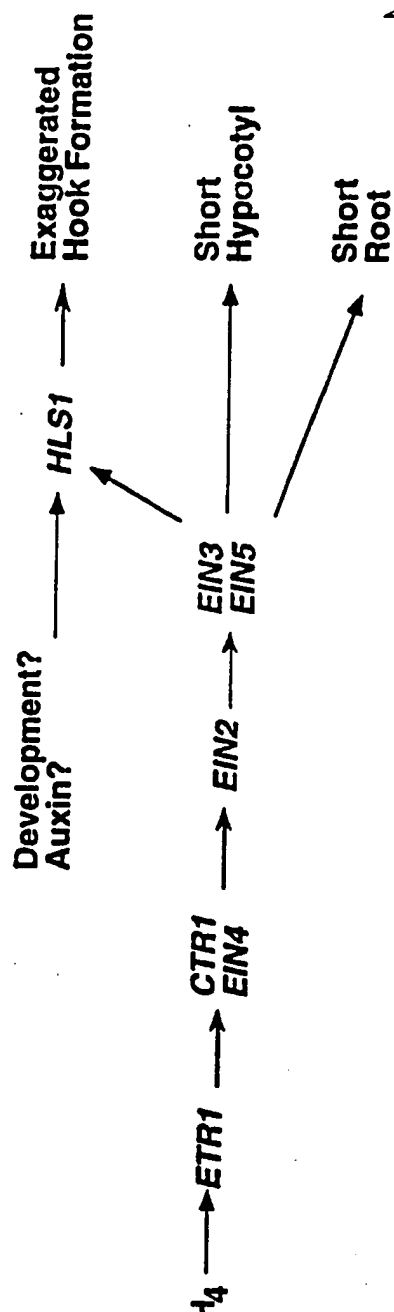


FIGURE 17

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pNLEIN3Bg12

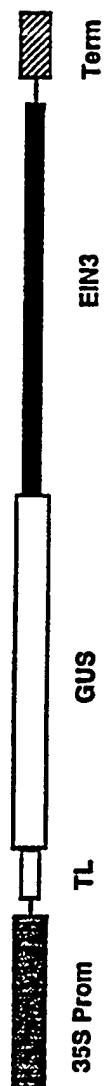


FIGURE 18

EIN3 cDNA

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TCTTCTTCTTCTTCTCCTCTTCCTCATCTCGTATCTCTAACTTTTGTCTGAAGTTCT
TTTGATGAAACTAGGGTTTATTATCTTCTCCTTCTTTTCCCATCACCATAGAA
AAGGCAGAGACCTTTTTCTTCATCATTTTTATTCTCCTTCTTCTTCTGCTGT
TCATTTCTCCAGGTTACAATGATGTTTAAATGAGATGGGAATGTGTGGAAACAT
GGATTTCTTCTTCTTCTGGATCACTTGGTGAAGTTGATTTCTGTCCTGTTCCACA
AGCTGAGCCTGATTCATTGTTGAAGATGACTATACTGATGATGAGATTGATG
TTGATGAATTGGAGAGGAGGATGTGGAGAGACAAAATGCGGCTTAAACGTCT
CAAGGAGCAGGATAAGGGTAAAGAAGGTGTTGATGCTGCTAAACAGAGGCA
GTCTCAAGAGCAAGCTAGGAGGAAGAAAATGTCTAGAGCTCAAGATGGGATC
TTGAAGTATATGTTGAAGATGATGGAAGTTTGTAAAGCTCAAGGCTTTGTTTAT
GGGATTATTCCGGAGAATGGGAAGCCTGTGACTGGTCTTCTGATAATTTAAG
GGAGTGGTGGAAAGATAAGGTTAGGTTTATCGTAATGGTCCTGCGGCTATTA
CCAAGTATCAAGCGGAGATAATATCCCGGGGATTCATGAAGGTAATAACCC
GATTGGACCGACTCCTCATACCTTGCAAGAGCTTCAAGACACGACTCTTGGA
TCGCTTTTGTCTGCGTTGATGCAACACTGTGATCCTCCTCAGAGACGTTTTCC
TTTGGAGAAAGGAGTTCTCCTCCGCGGTGGCCTAATGGGAAAGAGGATTGG
TGGCCTCAACTTGGTTTGCCTAAAGATCAAGGTCCTGCACCTTACAAGAAGC
CTCATGATTTGAAGAAGGCGTGGAAGTGGCGCTTTTACTGCGGTTATCAA
GCATATGTTTCTGATATTGCTAAGATCCGTAAGCTCGTGAGGCAATCTAAAT
GTTTGCAGGATAAGATGACTGCTAAAGAGAGTGCTACCTGGCTTGCTATTATT
AACCAAGAAGAGTCCTTGGCTAGAGAGCTTTATCCCGAGTCATGTCCACCTC
TTTCTCTGTCTGGTGGAAGTTGCTCGCTTCTGATGAATGATTGCAGTCAATAC
GATGTTGAAGGTTTTCGAGAAGGAGTCTCACTATGAAGTGGAAGAGCTCAAGC
CAGAAAAAGTTATGAATTTCTCAAACCTTTGGGATGGTTGCTAAAATGCATGAC
TTTCTGTCAAAGAAGAAGTCCAGCAGGAACTCGGAATTCATGAGAAAGA
GAAAGCCAAACAGAGATCTGAACACTATTATGGACAGAACCGTTTTCACCTG
CGAGAATCTTGGGTGTGCGCACAGCGAAATCAGCCGGGGATTCTGGATAG
GAATTCGAGAGACAACCATCAACTGGCATGTCCACATCGAGACAGTCGCTTA
CCGTATGGAGCAGCACCATCCAGGTTTCATGTCAATGAAGTTAAGCCTG
TAGTTGGATTTCTCAGCCAAGGCCAGTGAACCTCAGTAGCCCAACCAATTGA
CTTAACGGGTATAGTTCTGAAGATGGACAGAAGATGATCTCAGAGCTCATG
TCCATGTACGACAGAAATGTCCAGAGCAACCAACCTCTATGGTCATGGAAA
ATCAAAGCGTGTCACTGCTTCAACCCACAGTCCATAACCATCAAGAACATCT
CCAGTTCCCAGGAAACATGGTGGAAGGAAGTTTCTTTGAAGACTTGAACATC
CCAAACAGAGCAAACAACAACAGCAGCAACAATCAAACGTTTTTTCAAG
GGAACAACAACAACAACATGTGTTTAAAGTTGACACTGCAGATCACAACAA
CTTTGAAGCTGCACATAACAACAATAACAGTAGCGGCAACAGGTTCCAG
CTTGTGTTTGATTCCACACCGTTTCGACATGGCGTCATTGATTACAGAGATGA
TATGTCGATGCCAGGAGTAGTAGGAACGATGGATGGAATGCAGCAGAAGCA
GCAGATGTATCCATATGGTTCTAAAGTCTTGGTAGTAGATTTCATCTTCTCTT
ATTTTATCTTTGTGTTCTTACATTCACTCAACCATGTAATATTTTCTCTGGG
TCTCTCTGTCTCTATCGCTTGTATGATGTGTCTGTAAGAGTCTCTAAAACTC
TCTGTTACTGTGTGCTTTGTCTCGGCTTGGTGAATCTCTCTGTCTCATCATCAG
CTTTAGTTACACACCCGACTTGGGGATGAACGAACACTAAATGTAAGTTTC
A

FIGURE 19A

EIN3 genomic

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AGAGCAGTGAGTATTNCCACNAGCCGCTTTGTTAATTACATATTAATTGTGTA
ATAATAATAATAAATGATGTCTTAAATTTTATGTGTAAGAAATGAAATTTAAATG
ATATATATGTATATTATATATCTANACATATATATATATATAAATAGAGTATAT
ATACTATGATCTATCTTCCTGATCTACAGAGAGACTCCACAAAGAAACGCAAA
TAAACAAAAGTCGCTTTCTAGCCACGTGATCTTTCGTCGACTTTTCTTCTTCTT
CTTCTTCTTCTCCTCTTCCTCATCTCGTATCTCTAACTTTTGTCTGAAGTTCTTTTG
ATGAAACTAGGGTTTATTATCTTCTCCTTCTTTTCCCATCACCATAGAAAAGG
CAGAGACCTTTTCTTCATCATTTTTATTCTCCTTCTTCTTCTGCTGTTCAATTC
TCCAGGTACTATACGCTTCTTCTTCTATTGATTTTTTAGGGTTATTATTGATACT
GAAGATGATGATAGGTTTATTCATAGGGTTTTACTAGATCGATGGTTTTACTTT
AGTTTACTAGTGTTTACACGATCTAATTTTCATGAGTTTATNCTACTTTTAGTTTT
TINNTTGGGTGAAGTTTTGTTTATTGTTTATAAATCGTTGATCTATTTGAAAATG
TTTTCTCTTCTTATTATATATGATCCTTTCTATATTTGGTTCCTATGTTGAAG
ATCTCATCCTTTTTTTTGGAAATTGAATCTGTTGATAATTTTTATTATCCGATTGA
TTATTTAGTTTAGGAGTGATTAAAATACGATCTGATTATGTGTTTATTACTTAAA
ACTTTGATTGAATTCGAAAAGCCCCCTTTTTTATAATTTAGGGTTTGATGATTTT
TTTAGTAAGTTGTTTGATTGAGAAGAAATATAATTGTAAGTATTAGTTTGTGTTG
TGTATTTGATTTGTTACAGGTTACAATGATGTTTAAATGAGATGGGAATGTGTGG
AAACATGGATTTCTTCTCTTCTGGATCACTTGGTGAAGTTGATTTCTGTCCTGT
TCCACAAGCTGAGCCTGATTCCATTGTTGAAGATGACTATACTGATGATGAGA
TTGATGTTGATGAATTGGAGAGGAGGATGTGGAGAGACAAAATGCGGCTTAA
ACGTCTCAAGGAGCAGGATAAGGGTAAAGAAGGTGTTGATGCTGCTAAACAG
AGGCAGTCTCAAGAGCAAGCTAGGAGGAAGAAAATGTCTAGAGCTCAAGATG
GGATCTTGAAGTATATGTTGAAGATGATGGAAGTTTGAAAGCTCAAGGCTTT
GTTTATGGGATTATTCCGGAGAATGGGAAGCCTGTGACTGGTGTCTTGATAA
TTTAAGGGAGTGGTGGAAAGATAAGGTTAGGTTTGATCGTAATGGTCTCGCGG
CTATTACCAAGTATCAAGCGGAGAATAATATCCCGGGGATTATGAAGGTAAT
AACCCGATTGGACCGACTCCTCATACCTTGCAAGAGCTTCAAGACACGACT
CTTGGATCGCTTTTGTCTGCGTTGATGCAACACTGTGATCCTCCTCAGAGAC
GTTTTCTTTGGAGAAAGGAGTTCCTCCTCCGTGGTGGCCTAATGGGAAGA
GGATTGGTGGCCTCAACTTGGTTTGCCTAAAGATCAAGGTCTGCACCTTAC
AAGAAGCCTCATGATTTGAAGAAGGCGTGGAAGTCGGCGTTTTGACTGCGG
TTATCAAGCATATGTTTCTGATATTGCTAAGATCCGTAAGCTCGTGAGGCAA
TCTAAATGTTTGCAGGATAAGATGACTGCTAAAGAGAGTGCTACCTGGCTTGC
TATTATTAACCAAGAAGAGTCCTTGGCTAGAGAGCTTTATCCCGAGTCATGTC

FIGURE 19B

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EIN3 peptide

MMFNEMGMCGNMDFSSGSLGEVDFCPVQAEPDSIVEDDYTDDEIDVDELE
RRMWRDKMRLKRLKEQDKGKEGVDAKQRQSSEQARRKKMSRAQDGILKYM
LKMMEVCKAQGFVYGIIPENGKPVGTASDNLREWWKDKVRFDNRNGPAAITKYQ
AENNIPGIHEGNNPIGPTPHTLQELQDTTLGSLLSALMQHCDPPQRRFPLEKGV
PPPWWPNGKEDWWPQLGLPKDQGPAPYKKPHDLKKAWKVGVLTAVIKHMFP
DIAKIRKLVQRQSKCLQDKMTAKESATWLAIINQEESLARELYPESCPPLSLSGG
SCSLLMNDCSQYDVEGFEEKESHYEVEELKPEKVMNSSNFGMVAKMHDFPVK
EEVPAGNSEFMRKRKPNRDLNTIMDRTVFTCENLGCAHSEISRGFLDRNSRDN
HQLACPHRDSRLPYGAAPSRFHVNEVKPVVGFPQPRPVNSVAQPIDLTGIVPE
DGQKMISELMSEMYDRNVQSNQTSVMENQSVSLLQPTVHNHQEHLQFFGN
MVEGSFFEDLNIPNRANNNNNSSNNQTFQGNNNNNNVFKFDTADHNNFEAAH
NNNNNSSGNRFQLVFDSTPFDMA SFDYRDDMSMPGVVGTMDGMQQKQQDV
SIWF

FIGURE 19C

EIL1 cDNA

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GGCCGCTTCAAACCTCTACAAACCCAGAAACCACCACACAGTAATTAATGTCT
CTTTCTTTCTTCCCATGTGATCTTTAACAGACTTTTCTTCTTATTCTCCATCTC
TGAAGTGTGGGGATTTCATCAAGACTTCCTTATCTGTTTCTTTATAAAACAA
GAGAGAGATACCACTTTTGGTGTCTTTATTTGCAACTCTTTCAGGTTAAAGA
AATCGATAGGCTCTGTTCTTGATTGTGGTGGAAGAGAcATGATGATGTTTAc
GAGATGGGAATGTATGGAAACATGGATTTCTTCTCTCCTCCACATCTCTCGA
tGTGtGtccATTACCACAAGCTGAACAAGAACCTGTagtTGAagaTGTCGACTACA
CCGATGATGAGATGGATGAGCTTGAGCAGAGGATGTGGAGAGACAAAATGC
GTTTGAAACGTCTCAAGGAGCAACAGAGTAAGTGTAAAGGAGGCGTCGATg
GTTCGAAACAGAGGCAGTcgCaAGAGCAAGCTAGGAGGAAGAAAAgtCTAGA
GCCCAAGATGGGATCTTGAAGTATATGTTGAAGATGAIGGAAGTTTGTAAG
CTCAAGGCTTTGTTTATGGTATTATTCCTGAGAAGGGTAAGCCTGTGACTGG
tGCTTCGGATaATTTGAGGGAATGGTgGAAAGATAAGGTTAGGTTTGATCGTA
ATGGTCCAgCTGCTATTGCTAAGTATCAGtCAGAGAATaATATTTCTGGAGGG
AGTAATGATTGTAAACAGCTTGgTTGGTCCAACACcgATACGcTTCAGGAGCT
TCAGGACACGACTCTTGgTTCgCTTTATCGGCTTTGATGCAACATTGTGAT
CCACCGCAGAGACGGTTTCCTTTGgaGAAaGGAGTTTCTcCACCTTGGTGGC
CTAATGGGAATGAAGAgTgGTGGccTcaGCTtGtTTACCAAATGAGCAAGGTCC
TCCTCTTATAAGAAGCCTCATGATTTGAAGAAAGCTTGGAAAgTCGGTGTtT
TaACTGCGGTGATCAAGCATATgTCGCGGATATTGCGAAGATCCGTAAGCT
TGTGAGGCAATCAAAATGCTTgCAGGATAAGATGACGGCGAAAGAGAGTGC
TACTTGGCTTGCCATTATTAACCAAGAAGAGGTTGTGGCTCGGGAgCTTTAT
CCCGAGTCATGCCCTCCTCTTTCTTCTTCTTCATCATTAGGAAGCGGGTCCG
TtcCATTAAATGATTGTAGCGAGTATGACGTTGaAGGTTTCGAGAAGGaaCaA
CATGGTTTCGATGTGGaAGAGCGGAAACCAGAGATAGTGATGATgCATCCTC
TAgCAAGCTTTGGGGTTgCTAAAATGCAACATTTTCCcATAAAGGAGGAGGT
CgCCAaCACGGTAAACTTAGAGTTCACGAGAAAGAGGAAGCAGAACAATGAT
ATGAATGTTATGGTAATGGACAGATCAGcAGGTTACACtGTGAGaATGGTca
GTGTCTCACAGCAAAATGAaTCTTGgATTTCAAGACAGGAGTTCAAGGGAC
AACCACCAGATgGTTTGTCCATATAGAGACAATCGTTTAgCGTATGGAGCAT
CCAAGTTTcATATGGGTGGAAIGAACTAGTAGTTCTCAGCAAcCAGTCCaa
CCGATCGAcATCGGGCGTTGGAGTTCCGGAAAACGGGCaGAAGATGAT
CACCGAGCTTATGGCCATGTACGACAGAAATGTCCAAAGCAACCAACGCC
TCCTACTTTGATGGAAAACCAAGCATGGTCATTGATGCAAAAGCAGCTCAG
AATCAGCAGCTGAATTTCAACAGTGGCAATCAAATGTTTATGCAACAAGGGA
CGAACAACGGGGTTAACAATCGGTTCCAGATGGTGTtTGATTGACACCATT
CGATATGGCAGCATTGATTACAGAGATGATTGGCAAACCGGAGCAATGGA
AGGAATGGGGAAGCAGCAGCAGCAGCAGCAGCAGCAGCAGCaAGATGTATCA
ATATGGTCTGAATATTACACAATCTGTGAATATTCACTCTTTCATAATAACT
CTGTTACCTACTTACCTGACTTGGGTATGTATTCTATTGCACCAACACTCAT
CTATATTGTTGATGATGATGAAGCCATCTATTTTTTTTTTGTGTCTGAAAGTC
ATTTAACTCGCTTCATTGTTTTAATAATGTCACTATCCATTGAACATCATTCTC
ATGCTACAAGTTTGATTCTTTGAGGCGGCCGC

FIGURE 20A

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EIL1 peptide

MMMFNEMGMYGNMDFSSSTS LDVCLPQAEQEPVVEDVDYTDDEMDVDE
LEKRMWRDKMRLKRLKEQQSKCKEGVDGSKQRQSQEQARRKKMSRAQDGIL
KYMLKMMEVCKAQQGFVYGIIPEKGKPVGTGASDNLREWWKDKVRFDNRNGPAAIA
KYQSENNISGGSNDCNSLVGPTPHTLQELQDITLGSLLSALMQHCDPPQRRF
PLEKGVSPPPWWPNGNEEWWPQLGLPNEQGPPPYKKPHDLKKA WKVGLTAV
IKHMSPDIKIRKLVRSKCLQDKMTAKESATWLAIINQEEVVARELYPESCPPL
SSSSSLGSGSLLINDCSEYDVEGFEEQHGFDVEERKPEIVMMHPLASFGVA
KMQHFPIKEEVATTVNLEFTRKRKQNNDMNVMVMDRSAGYTCENGQCPSHKM
NLGFQDRSSRDNHQMVCPYRDNR LAYGASKFHMGGMKLVVPQQPVQPIDLS
GVGVPENGQKMITELMAMYDRNVQSNQTPPTLMENQSMVIDAKAAQNQQLNF
NSGNQMFMQQGTNNGVNNRFQMVFDSTPFDMAAFDYRDDWQTGAMEGMGK
QQQQQQQQQDVSIWF

FIGURE 20B

EIL2 cDNA

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CAGATTCTATGGATATGTATAACAACAATATAGGGATGTTCCGGAGTTTAGTTT
GTAGCTCGGCGCCTCCATTTACAGAGGGACATATGTGTTCTGATTTCGCATAC
GGCTTTGTGCGATGATCTGAGTAGTGATGAGGAAATGGAAATAGAGGAGCTT
GAGAAGAAGATCTGGAGAGACAAGCAGCGTTTAAAGCGGCTCAAGGAAATG
GCGAAGAACGGTCTAGGAACAAGATTGTTGTTGAAGCAGCAACATGATGATT
TTCCAGAGCACTCTAGTAAGAGAACCATGTACAAGGCACAAGATGGGATCTT
GAAGTACATGTCTGAAGACAATGGAGCGATATAAAGCTCAAGGTTTTGTTTATG
GGATTGTGTTAGAGAATGGGAAAACGGTAGCGGGATCTTCTGATAATCTCCG
TGAATGGTGGAAAGACAAAGTGAGGTTTGATAGGAACGGCCCAGCTGCTATA
ATCAAGCACCAAAGGGATATCAATCTTCTGATGGAAGTGATTTCAGGGTCTGA
GGTTGGGGATTCTACCGCACAGAAGTTGCTTGAGCTTCAAGATACTACTCTT
GGAGCTCTGTTATCGGCTCTGTTTCTCTCACTGCAACCCTCCTCAGAGGCGGT
TTCCGTTGGAGAAAGGCGTGACACCGCCATGGTGGCCAACGGGGAAAGAAG
ATTGGTGGGATCAACTGTCTTTACCGGTTGATTTTCGAGGTGTTCCGCCACCT
TACAAGAAGCCTCATGATCTCAAGAAGCTGTGGAAAATTGGTGTGTTGATTGG
TGTAATCAGACATATGGCTTCTGACATTAGCAACATACCCAATCTCGTGAGAC
GGTCTAGAAGTTTGCAGGAGAAAATGACGTCAAGAGAAGGCGC
TTTATGGCTCGCTGCTCTTTACCGAGAAAAGGCTATTGTTGATCAAATAGCCA
TGTCTAGAGAAAACAACAACACTTCTAACTTTCTTGTTCCTGCAACCGGTGGA
GACCCAGATGTTTTGTTTCCTGAATCTACAGACTATGATGTTGAACTGATTGG
TGGCACTCATCGGACCAATCAGCAGTATCCTGAATTTGAAAACAACATAAC
TGTGTTTACAAGAGAAAAGTTTGAAGAAGATTTGGGATGCCAATGCATCCAAC
ACTCCTAACATGTGAGAACAGTCTCTGTCTTATAGCCAACCACATATGGGA
TTTCTTGACAGGAACTTAAGAGAGAATCACCAAATGACTTGTCTTATAAAGT
CACTTCCTTCTACCAACCAACTAAACCCTATGGTATGACGGGTTTAAATGGTTC
CTTGTCGGGATTATAACGGGATGCAGCAGCAGGTTTCAGAGCTTTCAAGACCA
GTTTAAATCATCCAACGATCTCTACAGACCAAAAGCTCCACAAAGAGGCAAC
GATGACTTGGTTGAGGATTTGAATCCTTCTCCTTCGACGCTGAATCAGAATCT
TGGTTTAGTCTTACCTACTGACTTCAATGGAGGTGAGGAAACAGTAGGAACA
GAGAACAATCTGCATAATCAAGGGCAAGAGTTGCCACATCTTGGATTCACT
AAAGAAAGCTTCAGAGTTTTCTTTTTATGTTTTCTAGTCTTTATAGCTTTGTCTC
TTGCTTATTCTCTCATTAAACACAGTTTTTGATCTCTCCATTTATAGCCCATG
TAGCAATGGAGAAGATTAGGTTTCATAATAAGTTAATAACCAAATTCAAA

FIGURE 21A

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EIL2 peptide

DSMDMYNNNIGMFRSLVCSSAPPFTEGHMCSDSHTALCDDLSSDEEMEIEEL
EKKIWRDKQRLKRLKEMAKNGLGTRLLKQQHDDFPEHSSKRTMYKAQDGILK
YMSKT MERYKAQGFVYGIVLENGKTVAGSSDNLREWWKDKVRFDRNGPAAIK
HQRDINLSDGSDSGSEVGDSTAOKLLELQDITLGALLSALFPHCNPPQRRFPL
EKGVTTPWWPTGKEDWWDQLSLPVDFRGVPPPYKKPHDLKKLWKIGVLIGVIR
HMASDISNIPNLVRRSRSLQEKMSTRREGALWLAALYREKAIVDQIAMSRENNNT
SNFLVPATGGDPDVLFPESDYDVELIGGTHRTNQQYPEFENNYNCVYKRKFE
EDFGMPMHPTLLTCENSLCPYSQPHMGFLDRNLRENHQMTCPYKVTSFYQPT
KPYGMTGLMVPCPDYNGMQQQVQSFQDQFNHPNDLYRPAKAPQRGNDDLVED
LNPSPTLNQNLGLVLPDFTNGGEETVGTENNLHNQGQELPTSWIQ

FIGURE 21B

EIL3 cDNA

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TTCCCCTGAGAACGACAGGAGAAAGAATAAAAACCCTAAATTTCTTTAATTTT
GGCGCTTCAGATTATCGTTGTTAAAGGTTTTTGATTGATTTTGTAAATGGGC
GATCTTGCTATGTCCGTAGCAGACATCAGGATGGAGAATGAGCCTGATGATT
TAGCTAGTGATAATGTTGCTGAGATTGATGTGAGTGATGAAGAGATTGATGCT
GACGACCTTGAGAGACGGATGTGGAAAGATCGTGTCAAGGCTTAAAGAATCA
AAGAGCGACAAAAAGCTGGCTCTCAAGGAGCTCAAAACGAAGGGAGACACC
TAAGAAAATCTCTGATCAAGCTCAGAGGAAGAAAATGTCTTAGAGCTCAAGAT
GGTATCCTTAAGTACATTGTTGAAGCTTATGGAAGTCTGCAAAGTTCGCGGGT
TTGTCTATGGTATAATACCGGAAAAGGGCAAGCCTGTGAGTTGGCTCCTCTG
ACAATATAAGAGCTTGGTGGAAAGAGAAAGTGAAGTTTGATAAGAA₂CGGTCT
GCTGCTATTGCTAAATACGAAGAGGAGTGTTTAGCGTTTGGGAAATCTGATGG
GAATAGGAATTCACAGTTTGTCTCCAGGATTTGCAAGATGCTACTTTAGGGT
CTTTGTTATCTTCTTTGATGCAACATTGTGATCCTCCTCAAAGGAAGTATCCGT
TGGAGAAAGGGACGCCTCCGCCTTGGTGGCCAACGGGGGAATGAAGAATGGT
GGGTGAAACTCGGTCTGCCTAAAAGCCAGAGTCTCCTTACCGAAAACCTC
ATGATCTCAAGAAGATGTGGAAGGTTGGAGTTTAAACGGCAGTGATCAATCAT
ATGTTACCTGATATTGCAAAGATTAAGAGGCATGTTTCGTCAGTCGAAATGTTT
ACAGGACAAGATGACAGCTAAAGAGAGTGCGATTTGGTTGGCGGTTTTGAAC
CAAGAGGAATCTTTGATTACAGCAGCCTAGCAGTGACAATGGAACTCCAATG
TGACTGAGACACATCGTAGGGGTAAATAACGCTGACAGGAGGAAACCTGTGGT
CAACAGTGACAGTGACTATGATGTTGATGGGACAGAGGAAGCTTCAGGTTCA
GTTTCATCTAAAGACAGTAGAAGAAATCAGATTCAAAAAGAACAACCAACAG
CCATCTCACATTCAGTAAGAGATCAAGATAAAGCAGAGAAACATCGCAGAAG
GAAAAGACCTCGAATTAGATCCGGAACGTGCAATCGACAAGAGGAAGAACA
CCTGAAGCTCAACAAAGAAACATCTTACCTGATATGAATCATGTTGATGCCC
CTCTGCTAGAATATAACATCAACGGTACTCATCAAGAGGACGATGTTGTCTGA
CCCAATATTGCCTTAGGACCAGAGGAT₂ATG₉TCTGGAAGTGTGGTTCCTG
AGITCAATA₂CC₂₂₂CATACTTATCTTCCACTTGTTAATGAACAACTATGATGC
CTGTAGACGA₂AGGCCAATGCTTTATGGACCCAAACCCTAACCAAGAGCT
TCAATTTGGGTCAAGGGTACAACCTTCTACAATCCCTCTGCAGTGTTTGTACATA
ACCAGGAAGACGACATTCTCCATACACAGATAGAAATGAATACACAAGCACC
ACCTCACACAGTGGGTTTCGAGGAGGCCCCAGGAGGAGTACTTCAACCCCT
TGGTTTACTCGGAAATGAAGACGGTGTAACAGGGAGTGAGTTGCCTCAGTAT
CAGAGTGGCATTCTGTCTCCATTGACTGACTTGGACTTTGACTATGGTGGTTT
TGGTGATGATTTCTCATGGTTTGGAGCTTAGTGTCTTGCCATTTTTTTGGGAG
ATTACATAGTTCAAAGGACATGGCAATAGTCTGGCTAGTACAGTTACTTTCT
CTTCTTCATTTCTTCTGATCTTATATTCTTCCTCTTTTTTCTTATAATATTTCT
TAGATTTGTTAAGAGAAACAATTTTCCTTTGAATAAGTTGCCAGAAGAACTGC
TTTGCCCGTTGTAATGGTCTCTAGGGAAAGCAGTTAGCGTATCATCATTTGTA
AATTTACCTGTGAG

FIGURE 22A

HLS1 cDNA:

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CTCCAACTTTTAAACTCATCATAAATAGTAAAAAAGTAGCCGGAAAAATAAA
ATAAAAAGTCTATTTCTCTTTCCTTTAAAATCCAAATCCTATAAACTCATAGCT
TTCTCTGTTCTTTACTTATACCTCACGTTATACATATATATAGAGTTTCTATA
AATGCTTCTCTTTCCTCTCGAACAAATCTTCCTCACTTCTCTCATTTCACAC
TCACCTTCCTCTCTATATATTAAACCCTATCTACTTAAGTCTTCTTCTAACTCT
AATCTCTCTCTCTATTTACTCTGCTTCTGTTCTCACTCTGAAAGAACCAAAAC
ATGACGGTGGTTAGAGAGTACGACCCGACCCGAGACTTAGTCGGCGTGGAG
GACGTGGAACGACGGTGTGAAGTCGGACCAAGCGGCAAGCTTTCTCTTTTCA
CCGACCTTTTGGGTGACCCGATTTGTAGAATCCGACATTCACCTTCCTATCT
CATGCTGGTGGCTGAGATGGGTACGGAGAAGAAGGAGATAGTGGGCATGATT
AGAGGATGTATCAAAACCGTTACATGTGGCCAAAACTCGATTTAAATCACAA
ATCTCAAAACGATGTCGTTAAGCCTCTTTACACTAAACTCGCTTACGTCTTGG
GCCTTCGCGTCTCTCCTTTTACAGGAGACAAGGGATTGGGTTTAAGCTCGT
GAAGATGATGGAGGAATGGTTTAGACAAAACGGAGCTGAGTATTCGTATATTG
CAACTGAGAACGATAATCAAGCTTCTGTGAATTTGTTACCGGGAAATGTGGT
TATTCGGAGTTTCGTACACCGTCGATTTTGGTTAACCCGGTTTACGCTCATCG
AGTTAATGTTTCGCGGCGAGTCACGGTTATCAAGTTAGAGCCGGTTGATGCT
GAGACGTTGTACCGAATCCGGTTTAGCACAACAGAGTTTTTCCGCGGGATA
TTGATTCGGTACTTAATAACAACTCTCGCTTGGGACTTTTCGTCTCGCGGTGCCA
CGTGGAAGCTGTTATGGATCCGGGTCTGGATCATGGCCCGGTTCCGGCTAAAT
TCCTCGAATATCCACCCGAGTCATGGGCCGTATTAAGCGTGTGGAATTGTAA
AGACTCGTTTCTGTTAGAAGTACGTGGAGCGTCGAGATTGAGACGTGTGGTG
GCTAAAACGACGCGAGTAGTTGATAAAACGTTGCCGTTTCTGAAACTACCTT
CGATACCGTCCGTTTTTGAACCTTTTGGACTTCATTTTATGTATGGAATCGGA
GGAGAAGGTCCACGCGCGGTGAAGATGGTGAAATCCTTGTGTGCTCACGCG
CATAACTTGGCTAAGGCAGGTGGTTGTGGTGTCTGTCGGCGGCGGAAGTTGCC
GGAGAAGACCCGTTGCGGCGAGGAATACCACATTGGAAAGTGCTATCGTGT
GACGAGGATCTTTGGTGTATAAAGCGGCTTGGAGATGACTATAGTGATGGTGT
TGTTGGTGAATTGGACTAAATCGCCACCTGGCGTTTCCATTTTGTAGACCCT
AGAGAATTTTAAACCTTTTTTTTAACTCTATAATATATATTCTCTATTAACCACT
TGATGTTAAATTAGGGGTTTTCTTCTAAGTTTATAGATTTTCTTGTTTTAGAATTA
ATCTTTTTTTAGGTAACCTTTTTTGCTTTTGTGTTTGTGTTTGTGTTTGTGG
GTGTTATAAATTA

FIGURE 23A

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CGTGGAAGCTGTTATGGATCCGGGTCTGGATCATGGCCCGGTTCCGGCTAAAT
TCCTCGAATATCCACCCGAGTCATGGGCCGTATTAAGCGTGTGGAATTGTAA
AGACTCGTTTTCTGTTAGAAGTACGTGGAGCGTCGAGATTGAGACGTGTGGTG
GCTAAAACGACGCGAGTAGTTGATAAACGTTGCCGTTTCTGAAACTACCTT
CGATACCGTCCGTTTTGAAACCTTTTGGACTTCATTTTATGTATGGAATCGGA
GGAGAAGGTCCACGCGCGGTGAAGATGGTGAAATCCTTGTGTGCTCACGCG
CATAACTTGGCTAAGGCAGGTGGTTGTGGTGTCTGGCGGCGGAAGTTGCC
GGAGAAGACCCGTTGCGGCGAGGAATACCACATTGGAAAGTGCTATCGTGT
GACGAGGATCTTTGGTGTATAAAGCGGCTTGGAGATGACTATAGTGATGGTGT
TGTTGGTGATTGGACTAAATCGCCACCTGGCGTTTCCATTTTGTAGACCCTA
GAGAATTTTAAACCTTTTTTTAACTCTATAATATATTTCTCTATTAACCACTT
GATGTTAAATTAGGGGTTTTCTTCTAAGTTTATAGATTTTCTTGTTTTAGAATTA
ATCTTTTTTTTAGGTAACTTTTTTGCTTTTTGTTTTGTTTTGTTTTGTTTGTGG
GTGTTATAAATTAgtgtaagaggtaatatctcctacttttgggttgtgtcttcttgttaaaggatctagc
ttttaagatacttttcttgtgccaacccaaaacgcgcacctgattattttccaagtagataaaatttcatgaac
gcactgatactataatgatgcaatttgtgtaagacgatacttggagataaaattacaatatgacaatgataga
aaatgttaccaataacgattagcattatcgtgtgtgccatcaagtataactaagagaaagacgcacattttcttta
agagtaaataaaaatatt

FIGURE 23B

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HLS1 polypeptide:

MTVVREYDPTRDLVGVEDVERRCEVGPSGKLSLFTDLLGDPICRIRHSPSYML
VAEMGTEKKEIVGMIRGCIKTVTCGQKLDLNHKSQNDVVKPLYTKLAYVLGLRV
SPFHRRQGIGFKLVKMMEEWFRONGAEYSYIATENDNQASVNLFTGKCGYSE
FRTPSILVNPVYAHRVNVSRRVTVIKLEPVDAETLYRIRFSTTEFFPRDIDSVLNN
KLSLGTFFVAVPRGSCYGS GSGSWPGSAKFLEYPPESWAVLSVWNCKDSFLL
EVRGASRLRRVAKTRRVVDKTLPLFLPSIPSVFEPFGLHFMYGIGGEGPRA
VKMVKSLCAHAHNLA KAGGCGVAAEVAGEDPLRRGIPHWKVLSCDEDLWCI
KRLGDDYSDGVVG DWTCHLAFPFLL

FIGURE 23C

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US95/07744

A. CLASSIFICATION OF SUBJECT MATTER

IPC(6) : C07K 14/415; C12N 5/00, 15/29; A01H 5/00, 7/00

US CL : 536/23.6, 23.1; 530/370; 800/200; 435/240 .4

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 536/23.6, 23.1; 530/370; 800/200

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

APS, GenEMBL sequence databases

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	Science, Volume 241, issued 26 August 1988, A. B. Bleecker et al, "Insensitivity to ethylene conferred by a dominant mutation in Arabidopsis thaliana", pages 1086-1089, see entire document.	1-17
A	Cell, Volume 72, issued 12 February 1993, J. J. Kieber et al, "CTR1, a negative regulator of the ethylene response pathway in Arabidopsis, encodes a member of the Raf family of protein kinases", pages 427-441, see entire document.	1-17
A	The Plant Cell, Volume 2, issued June 1990, P. Guzman et al, "Exploiting the triple response of Arabidopsis to identify ethylene-related mutants", pages 513-523, see entire document.	1-17



Further documents are listed in the continuation of Box C.



See patent family annex.

* Special categories of cited documents:	T	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
A document defining the general state of the art which is not considered to be of particular relevance	X	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
E earlier document published on or after the international filing date	Y	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
L document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	G	document member of the same patent family
O document referring to an oral disclosure, use, exhibition or other means		
P document published prior to the international filing date but later than the priority date claimed		

Date of the actual completion of the international search 14 SEPTEMBER 1995	Date of mailing of the international search report 05 OCT 1995
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Facsimile No. (703) 305-3230	Authorized officer <i>Elizabeth C. Kemmerer</i> ELIZABETH C. KEMMERER Telephone No. (703) 308-0196